(12) INTERNATIONAL APPLICATION PUBLISHED UNDER THE PATENT COOPERATION TREATY (PCT)

(19) World Intellectual Property Organization

International Bureau



(43) International Publication Date 6 January 2005 (06.01.2005)

PCT

(10) International Publication Number WO 2005/000335 A2

- (51) International Patent Classification⁷: 48/00, C12Q 1/68, A61P 39/00
- A61K 38/00,
- (21) International Application Number:

PCT/US2004/017322

- (22) International Filing Date:
 - 2 June 2004 (02.06.2004)
- (25) Filing Language:

English

(26) Publication Language:

English

(30) Priority Data:

60/474,606

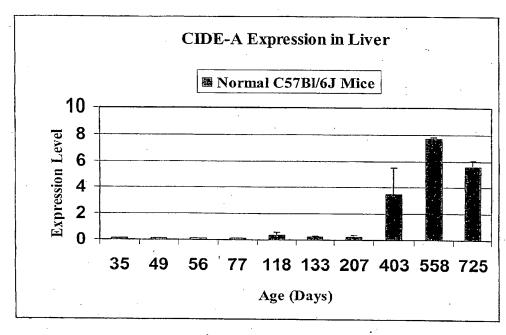
2 June 2003 (02.06.2003) US

- (71) Applicant (for all designated States except US): OHIO UNIVERSITY [US/US]; Technology Transfer Office, 20 East Circle Drive, Athens, 11 45701 (US).
- (72) Inventors; and
- (75) Inventors/Applicants (for US only): KOPCHICK, John, J. [US/US]; 4 Orchard Lane, Athens, 36 45701 (US). KELDER, Bruce [US/US]; 346 Carroll Road, Athens, Ohio 45701 (US). BOYCE, Keith S. [US/US]; 2589 Cole Road, Wexford, Pennsylvania 15090 (US). KRIETE, Andres [US/US]; 1222 Driftwood Drive, Pittsburgh, Pennsylvania 15243 (US).

- Agents: BROWDY AND NEIMARK, P.L.L.C. et al.; 624 Ninth Street N.W. II Suite 300, Washington, DC 20001-5303 (US).
- (81) Designated States (unless otherwise indicated, for every kind of national protection available): AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD. MG, MK, MN, MW, MX, MZ, NA, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY, TJ, TM. TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, **ZW**.
- (84) Designated States (unless otherwise indicated, for every kind of regional protection available): ARIPO (BW, GH, GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW), Eurasian (AM, AZ, BY, KG, KZ, MD, RU, TJ, TM), European (AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PL, PT, RO, SE, SI, SK, TR), OAPI (BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG).

[Continued on next page]

(54) Title: DIAGNOSIS AND TREATMENT METHODS RELATED TO AGING, ESPECIALLY OF LIVER



(57) Abstract: Mouse genes differentially expressed in comparisons of older and younger livers by gene chip analysis have been identified, as have corresponding human genes and proteins. The human molecules, or antagonists thereof, may be used for protection against faster-than-normal biological aging, or to achieve slower-than-normal biological aging. The human molecules may also be used as markers of biological aging.



MO Justile

WO 2005/000335 A2



Published:

 without international search report and to be republished upon receipt of that report For two-letter codes and other abbreviations, refer to the "Guidance Notes on Codes and Abbreviations" appearing at the begining of each regular issue of the PCT Gazette.

1

DIAGNOSIS AND TREATMENT METHODS RELATED TO AGING, ESPECIALLY OF LIVER

This application claims the benefit, under 35 USC 119(e), of U.S. Provisional application 60/474,606, filed June 2, 2003, which is hereby incorporated by reference in its entirety.

Cross-Reference to Related Applications

5

10

15

20

25

30

35

Anti-Aging Applications. Mice with a disrupted growth hormone receptor/binding protein gene enjoy an increased lifespan. In U.S. Prov. Appl. 60/485,222, filed July 8, 2003 (Kopchick8) mouse genes differentially expressed in comparisons of gene expression in growth hormone receptor/binding protein gene-disrupted mouse livers and normal mouse livers were identified, as were corresponding human genes and proteins. It was suggested that the human molecules, or antagonists thereof, could be used for protection against faster-than-normal biological aging, or to achieve slower-than-normal biological aging. It was also taught that the human molecules may also be used as markers of biological aging.

In provisional application Ser. No. 60/566,068, filed April 29, 2004 (our docket Kopchick14-USA), our research group used a gene chip to study the genetic changes in the muscle of C57Bl/6 mice that occur at various intervals of the aging process. Differential hybridization techniques were used to identify mouse genes that are differentially expressed in mice, depending upon their age. The level of gene expression of approximately 10,000 mouse genes (from the Amersham Codelink UniSet Mouse I Bioarray, product code: 300013) in the muscle of mice with average ages of 35, 49, 77, 118, 133, 207, 403, 558 and 725 days was determined. In essence, complementary RNA derived from mice of different ages was screened for hybridization with oligonucleotide probes each specific to a particular mouse gene, each gene

2

in turn representative of a particular mouse gene cluster (Unigene). Mouse genes which were differentially expressed (younger vs. older), as measured by different levels of hybridization of the respective cRNA samples with the particular probe corresponding to that mouse gene, were identified. Related human genes and proteins were identified by sequence comparisons to the mouse gene or protein.

5

25

30

35

Anti-Diabetes Applications. In U.S. Provisional Appl. 10 Ser. No. 60/458,398 (our docket Kelder1-USA), filed March 31, 2003, members of our research group describe the identification of genes differentially expressed in normal vs. hyperinsulinemic, hyperinsulinemic vs. type II diabetic, 15 or normal vs. type II diabetic mouse liver. Forward- and reverse-substracted cDNA libraries were prepared, clones were isolated, and differentially expressed cDNA inserts were sequenced and compared with sequences in publicly available sequence databases. The corresponding mouse and 20 human genes and proteins were identified.

The purpose of our research group's provisional application Ser. No. 60/460,415 (our docket: Kopchick6-USA), filed April 7, 2003, was similar, but complementary RNA, derived from RNA of mouse liver, was screened against a mouse gene chip. See also 60/506,716, filed Sept. 30, 2003 (Kopchick6.1).

Gene chip analyses have also been used to identify genes differentially expressed in normal vs. hyperinsulinemic, hyperinsulinemic vs. type II diabetic, or normal vs. type II diabetic mouse pancreas, see U.S. Provisional Appl. 60/517,376, filed Nov. 6, 2003 (Kopchick12) and muscle, see U.S Provisional Appl. 60/547,512, filed Feb. 26, 2004 (Kopchick15).

Other differential hybridization applications. The use of differential hybridization to identify genes and proteins

20

25

30

is also described in our research group's Ser. No. PCT/US00/12145 (Kopchick 3A-PCT), Ser. No. PCT/US00/12366 (Kopchick4A-PCT), and Ser. No. 60/400,052 (Kopchick5).

5 All of the foregoing applications are hereby incorporated by reference in their entirety.

BACKGROUND OF THE INVENTION

Field of the Invention

10 The invention relates to various nucleic acid molecules and proteins, and their use in (1) diagnosing aging, or adverse conditions associated with the aging process, and (2) protecting mammals (including humans) against the aging process or adverse conditions associated with the aging 15 process.

Description of the Background Art

The mechanisms that cause aging (the decline in survival and reproductive ability with advancing age) have puzzled our society and scientific community for centuries. The two major theories center on the question of whether normal aging is an evolutionarily-genetically preprogrammed pathway of internal changes or is a normal consequence of existence where there is an accumulation of molecular and cellular damages. Hypotheses of such accumulated damage include free radical-oxidative damage, defective mitochondria, somatic mutations, progressive shortening of telomeres, programmed cell death, impaired cell proliferation and numerous others (1). The current belief is that aging is not a programmed process in that, to date, no genes are known to have evolved specifically to cause damage and aging. The one factor that has been shown to extend the lifespan in organisms from yeast to mice has been a reduction in caloric intake (2, 3). Recent data suggests that caloric restriction may also be relevant for primates, 35 . including humans (4-6). Unfortunately, it is unlikely that

4

most people will be able to maintain the strict dietary control required to reap the benefits of this finding. Therefore, since the mechanism(s) by which caloric restriction extends lifespan are unknown, the elucidation of such mechanisms could lead to the development of alternative strategies to yield similar benefits.

5

10

15

20

25

30

35

Numerous groups are presently engaged in identifying genes and pathways that are involved in the aging process. A growing list of genes that extend adult longevity have been identified and a large proportion of these genes are involved with hormonal signals. Many of these genes and the corresponding endocrine systems are conserved among a wide variety of eukaryotes. What is becoming clear, at least in lower animal species, is that those pathways that provide advantages to development and growth early in life may impart negative consequences in later life. The clearest example of a genetic pathway affecting adult lifespan has been described in the nematode, Caenorhabditis elegans. When food is abundant, C. elegans develops directly to the reproductive adult through four larval stages in three days. Under adverse conditions such as caloric restriction or high population density, C. elegans enters the Dauer diapause, a non-feeding, stress-resistant larval state. Genetic analysis has identified that mutation of single genes involved in dauer formation (Daf) greatly extend the adult These genes involve the highly-conserved lifespan (7). insulin/IGF-like signal transduction pathway. Ligand binging to the daf-2 insulin-like receptor results in a kinase signaling cascade to phosphorylate the forkhead transcription factor, daf-16. This phosphorylation sequesters daf-16 to the cytoplasm and results in reproductive maturity and aging. In the absence of ligand and signal transduction, the unphosphorylated, daf-16 localizes to the nucleus and regulates the transcription of its target genes that promote dauer formation, stress resistance and extended longevity (8). A similar pathway

5

has been described in Drosophilia melanogaster. Mutation of the gene encoding insulin-like receptor (InR) or the gene encoding insulin-receptor substrate (chico) also extends the normal life-span (9,10). Vertebrate homologues of daf-16 down-regulate genes promoting cell progression, induce genes involved in DNA-damage repair and up-regulate genes that reduce intracellular reactive oxygen species (ROS) (11,12). A second C. elegans gene, clk-1, has also been linked to the reduction of ROS and an extended life-span. While the effect of daf-2 mutants result in a reduction of mitochondrial ROS, clk-1 mutants reduce extramitochondrially produced ROS. Since the majority of cellular ROS is produce in the mitochondria during the process of electron transport, it is not surprising that clk-1 mutants have only a moderately extended life-span. C. elegans containing daf-2/clk-1 double mutations, however, exhibit a very long lifespan (13).

5

10

15

20

25

30

35

Decreased IGF-1 signaling may also extend longevity in mice. Four mouse models with deficiencies in pituitary endocrine action have demonstrated retarded aging. In the *Prop1* and *Pit1* models, pituitary production of growth hormone (GH), prolactin (PRL) and thyroid stimulating hormone (TSH) are ablated. These mice have reduced growth rates, reduced adult body size and live 40 to 60% longer than normal mice (14,15). Unfortunately, it is not possible to determine which of the ablated hormones is responsible for the increased longevity of the models.

A more straightforward model was developed that targeted the deletion of the growth hormone receptor (GHR-KO) (16). This mouse line was derived from a founder animal by homologous recombination resulting in deletion and gene substitution of most of the fourth exon and part of the fourth intron of the GHR/BP gene. These mice also exhibit reduced body size and extended life-span and more directly implicates the GH/IGF-1 axis (17, 17a). Recently, evidence for a direct role of IGF-1 receptor signaling in affecting

6

the aging process was provided by the targeted disruption of the IGR-1 receptor (Igf1r) (18). Heterozygous females, but not males, possess 50% fewer receptors for IGF-1, live 33% longer than wild-type females and also display greater resistance to oxidative stress. Tyrosine phosphorylation of the intracellular signaling molecule, Shc, was also decreased in the Igf1r +/- females. Mice containing the targeted deletion of p66shc also have increased resistance to oxidative stress and a 30% increase in life span (19). While the IGF-1 axis appears to be involved in the aging process, the mechanism by which it does so remains unknown. However, these findings demonstrate that it is possible to identify specific genetic pathways that affect the aging process. The finding that caloric restriction of these mouse models can further extend their life-span suggests that multiple pathways exist that affect the aging process (20). Therefore, research to identify these pathways and the genes involved in the aging process is of great importance.

5

10

15

20

25

30

35

The role of growth hormone in aging is further discussed in Vance, ML, "Can Growth Hormone Prevent Aging," New Engl. J. Med., 348: 779-80 (Feb. 27, 2003).

Gene-Chip Based Identification of genes involved in aging of liver

Several groups have begun to utilize DNA microarrays to measure differences in gene expression caused by the aging process. However, these experiments are extremely limited in regards to the number of aging time points or experimental conditions.

Cao, S.X., et al., "Genomic profiling of short- and long-term caloric restriction effects in the liver of aging mice", Proc. Natl. Acad. Sci. USA, 98:10630-10635 (2001) used Affymetrix microarray technology to study the changes in expression levels of 11,000 genes in liver tissue of 7

7

month-old mice compared to 27 month-old mice. In this analysis, the expression of 20 genes increased at least 1.7fold with age while the expression of 26 genes decreased at least 1.7-fold with age. We have compared the differentially expressed genes described by Cao et al., to those that we have found to be differentially expressed using the Amersham platform. Of the 20 up-regulated genes, 10 had links from Affymetrix to Amersham through Unigene. Only one of Cao's up-regulated genes, Heat shock protein (L07577/NM_010410) was identified as differentially expressed in our analysis (increased 2.2-fold from weeks 2 to 4). Cao's 26 down-regulated genes, 10 had links from Affymetrix to Amersham through Unigene. Only one of these downregulated genes (Mouse TIS21 gene, M64292/NM 007570) was identified as differentially expressed in our analysis. However, we found the expression of this gene to increase 2.07-fold with age.

5

10

15

Tollet-Egnell, P., et al., "Gene expression profile of the aging process in rat liver: normalizing effects of growth hormone replacement, Mol. Endocrinol., 15(2):308-18 20 (2001) used microarray technology to study the effect of aging and growth hormone treatment on the expression of 3,000 different genes in the rat liver. The proteins which were over-expressed in the older rat were glucose-6phosphate isomerase (x1.8), pyruvate kinase (x4.8), hepatic 25 product spot 14 (2.4x), fatty acid synthase (1.9x), staryl CoA desaturase (1.7x), enoyl CoA hyydratase (1.7x), peroxisome proliferator activated receptor- α (1.7x), 3ketoacyl-CoA thiolase (1.7x), 3-keto-acyl-CoA peroxisomal thiolase (1.9x), CYP4A3 (3.3x), glycerol-3-phosphate 30 dehydrogenase (1.7x), NAPDH-cytochrome P450 oxidoreductase (4.7x). CUP2C7 (1.9x), CYP3A2 (2.8x), Δ -aminoevulinate synthase (2.3x). The under-expressed proteins were glucose-6-phosphatase (0.3x), farnesyl pyrophosphate synthase (0.5x), carnitine octanoyltransferase (0.5x), mitochrondrial 35 genome (16S ribosomal RNA)(0.3x), mitochondrial cytochrome c

oxidase II (0.4x), mitochondrial NADH dehydrogenase SU 5 (0.3x), mitochondrial cytochrome b (0.4x), mitochondrial NADH dhydrogenase SU 3 (0.5x), NADH-ubiquinone oxidoreductase (SU CI-SGDH and SU 39kDa) (both 0.5x), ubiquinol-cytochrome c reductase (Rieske iron-sulfur prote in and core 1) (both 0.5x), CYP2C12 (0.4x), cystathione γ -lyase (0.3x), biphenyl hydrolase-related protein (0.5x), glutathione S-transferase (class pi) (0.3x), α -1 macroglobulin (0.5x), BRAK related protein (0.3x), α -2u-globulin (0.4x), cAMP-dependent transcription factor mATF4 (0.5x), DAP-like kinase (0.5x), PCTAIRE-1 (0.5x), collagen α -1 (0.4x), histone H2A (0.5x), and S-100 protein α (0.5x).

Of the genes up-regulated in the older rat according to Tollet-Egnall, two have mouse cognates which we found to be up-regulated in the mouse liver. These were fatty acid synthase and stearyl CoA desaturase. A third, aminoevulinate synthase, has a mouse cognate which we found to be down-regulated in the older mouse. Two genes found by Tollet-Egnall to be down-regulated in the older rat were found by us to have cognates down-regulated in the older mouse: carnitine octanoyltransferase and CYP2C12.

See also Dozmorov I, Bartke A, Miller RA., "Array-based expression analysis of mouse liver genes: effect of age and of the longevity mutant Propldf", J. Gerontol., 56A: B52-57 (2001). Liver mRNA levels were measured in Ames dwarf mice (homozygous for the df allele at the Propl locus; live 40% to 70% longer than nonmutant siblings) and in control mice at ages 5, 13 and 22 months. "The analysis showed seven genes where the effects of age reach p < .01 in normal mice and six others with possible age effects in dwarf mice, but none of these met Bonferroni-adjusted significance thresholds. Thirteen genes showed possible effects of the df/df genotype at p < .01. One of these, insulin-like growth factor 1 (IGF-1), was statistically significant even after adjustment for multiple comparisons; and genes for two IGF-binding proteins, a cyclin, a heat shock protein, p38

C

mitogen-activated protein kinase, and an inducible cytochrome P450 were among those implicated by the survey. In young control mice, half of the expressed genes showed SDs that were more than 58% of the mean, and a simulation study showed that genes with this degree of interanimal variation would often produce false-positive findings when conclusions were based on ratio calculations alone (i.e., without formal significance testing). Many genes in our data set showed apparent young-to-old or normal-to-dwarf ratios above 2, but the large majority of these proved to be genes where high interanimal variation could create high ratios by chance alone, and only a few of the genes with large ratios achieved p < .05. The proportion of genes showing relatively large changes between 5 and 13 months, or from 13 to 22 months of age, was not diminished by the df/df genotype, providing no support for the idea that the dwarf mutation leads to global delay or deceleration of the pace of age-dependent changes in gene expression."

20

25

30

35

15

5

10

Gene-Chip Based Identification of Genes involved in aging of other organs and tissues

Gene expression profiling has been performed on skeletal muscle tissue of mice at 5 verses 30 months of age with or without caloric restriction (21). In this analysis, the expression of 113 genes was found to be changes by at least two-fold in 5-month old mice compared to 30-month old mice. Caloric restriction of comparable mice caused a reversal of the altered gene expression of 33 genes. Similar analyses have also been performed on mouse brain and heart (22,23).

Weindruch, et al., "Microarray profiling of gene expression in aging and its alteration by caloric restriction in mice" in Symposium: Calorie Restriction: effects on Body Composition, Insulin Signaling and Aging

10

918S-923S (2001)(21) compared expression in gastrocnemius muscle from 5- and 30-month old C57BL/6 mice, with and without caloric restriction. In this analysis, the expression of 113 genes was found to be changed by at least two-fold in 5-month old mice compared to 30-month old mice. Caloric restriction of comparable mice caused a reversal of the altered gene expression of 33 genes.

5

10

15

20

25

30

35

Of the 6347 genes surveyed in the oligonucleotide microarray, only 58 (0.9%) displayed a greater than 2 fold increase in gene expression as a function of aging, whereas 55(0.9%) displayed a greater than 2 fold decrease. Of the genes positively correlated with aging, 16% could be assigned to stress responses. The largest differential expression between young and aged animals (3.8 fold) was the mitochondrial sarcomeric creatine kinase.

Of the genes negatively correlated with aging, 13% were involved in energy metabolism. A noteworthy number were genes encoding biosynthetic enzymes (cytochrome P450 IIC12, squalene synthase, stearoyl-CoA desaturase, EF-1-gamma. Another down regulator was a CpG binding protein, MeCP2.

Weindruch further reported that age-related changes in gene expression profile were "remarkably attenuated" by caloric restriction.

What appears to be the same experiment is discussed in Lee, et al., "Gene expression profile of aging and its retardation by caloric restriction," Science, 285: 1390 (Aug. 27, 1999). This papers lists the individual genes which were differentially expressed by more than 2-fold, and classifies them as energy metabolism, neuronal factors, protein metabolism, stress response, biosynthesis, calcium metabolism or DNA repair genes.

Welle, et al., "Skeletal muscle gene expression profiles in 20-29 year old and 65-71 year old women," Exper. Gerontol., 39: 369-77 (2004) and available electronically as doi:10.1016/j.exger.2003.11.011 studied gene expression and physical condition in seven young and eight older women.

1.7

With respect to physical condition, the measured or calculated parameters were total body mass, lean body mass, left leg lean mass (by biopsy), maximum isometric left knee extension force, left knee extension force/left keg lean mass, Peak VO_2 /lean body mass, and Peak VO_2 /left leg lean mass.

5

10

15

20

25

30

35

There were 1178 "probe sets" (representing 1053 different Unigene clusters) for which differential expression was detected; 550 for which expression was higher in older women, and 628 the inverse effect. The differences ranged from 1.2 to 4 fold; most (78A%) were less than 1.5 fold. The complete list of differentially expressed genes is given in the Rochester Muscle database website, www.urmc.rochester.edu/smd/crc/swindex (".html" omitted, in accordance with USPTO requirements, so that the publication of this application will not create an active hyperlink).

The gene most highly overexpressed in older muscle was p21 (cyclin-dependent kinase inhibitor 1A) (4.01 fold). This one of several genes (see Welle Table 2) which are potentially related to DNA damage and repair. Welle also thought it noteworthy how many of the differentially expressed genes were ones that encode proteins which bind to pre-mRNAs or mRNAs (see Welle Table 3).

See also Lee et al., Science, 285 :1390-93 (1999) and Nature Genetics 25: 294-7 (2000) (bioarray study of changes in mouse cerebellum and neocortex to detect age-associated genes).

Non-Gene Chip Differential/Subtractive hybridization studies

The papers collected in this section deal principally with type II diabetes, which is an aging-related disease.

Sreekumar, et al., "Gene expression profile in skeletal muscle of type 2 diabetes and the effect of insulin treatment," *Diabetes* 51: 1913 (June 2002) surveyed 6,451 genes, and identified 85 genes for which there was an

12

alteration in skeletal muscle transcription in diabetic patients after withdrawal of insulin treatment. Subsequent insulin treatment resulted in further changes in transcription of 74 of the 85 genes (15 increased, 59 decreased), and also resulted in alteration of 29 additional gene transcripts.

5

10

15

20

25

Mootha, et al., "PCG-1 α responsive genes involved in oxidative phosphorylation are coordinatively downregulated in human diabetes," Nature Genetics 34(3); 267 (July 2003), used DNA microarrays to detect changes in the expression of sets of related genes, rather than of individual genes. They classified over 22,000 genes into 149 data sets; some of these data sets overlapped. They looked for a statistical correlation between the overall rank order of the genes in differential expression, and the groups to which the genes belonged. Expression was compared pairwise among three groups: males with normal glucose tolerance; males with impaired glucose tolerance; and males with type 2 diabetes. The set with the highest enrichment score (the one whose members ranked highly most often relative to chance expectation) was an internally curated set of 106 genes involved in oxidative phosphorylation. While the average decrease for the individual genes was modest (~20%), it was also consistent, being observed in 89% (94/106) of the genes in question. This paper is reviewed by Toye and Gauguier, "Genetics and functional genomics of type 2 diabetes mellitus", Genome Biology, 4: 241 (2003).

5

10

15

20

25

30

35

Patti, et al., "Coordinated reduction of genes of oxidative metabolism in humans with insulin resistance and diabetes: Potential role of PGC1 and NRF1", Proc. Nat. Acad. SCi. (USA), 100(14): 8466 (July 8, 2003) used microarrays to analyze skeletal muscle expression of genes in nondiabetic insulin-resistant subjects at high risk for diabetes (based on family hisotry of diabetes and Mexican-American ethnicity) and diabetic Mexican-American subjects. Of 7,129 sequences represented on the microarray, 187 were differentially expressed between control and diabetic subjects. However, no single gene remained significantly differentially expressed after controlling for multiple comparison false discovery by using the Benjamini-Hochberg method, see Benjamini, et al., J. R. Stat. Soc. Sert. B. 57:289-300 (1995); Dudait, et al., Stat. Sin. 12: 111-139 (2002). Consequently, Patti et al. sought to identify groups of related genes with similar patterns of differential expression using MAPP FINDER and ONTOEXPRESS. According to MAPP FINDER, the top-ranked cellular component terms were mitochondrion, mitochondrial membrane, mitochondrial inner membrane, and ribosome, and the topranked process term was ATP biosynthesis. According to ONTOEXPRESS, the over-represented groups were energy generation, protein biosynthesis/ribosomal proteins, RNA binding, ribosomal structural protein, and ATP synthase complex.

Huang, Xudong, "Identification of abnormally expressed genes in skeletal muscle contributing to insulin resistance and type 2 diabetes", Thesis, document id: 9576 Lunds University 2002, reported differential expression of the mitochondrially-encoded ND1 gene in human diabetic patients and of the nuclear-encoded cathepsin L gene in mice.

Standaert, et al., "Skeletal muscle insulin resistance in obesity-associated type 2 diabetes in monkeys is linked to a defect in insulin activation of protein kinase C-

14

zeta/lambda/iota Diabetes 51: 2936 (Oct. 2002). the authors concluded that defective activation of atypical PKCs played an important role in the pathogenesis of peripheral insulin resistance in both obese prediabetic and diabetic monkeys. They attributed this linkage to the apparent requirement for aPKCs during insulin-stimulated glucose transport.

5

10

15

20

25

30

Srommer, et al., Am. J. Physiol., "Skeletal muscle insulin resistance after trauma: insulin signaling and glucose transport", 275(2 Pt. 1): E3518(Aug. 1998) concluded that insulin resistance in skeletal muscle after surgical trauma is associated with reduced glucose transport but not with impaired glucose signaling to PI 3-kinase or its downstream target, Akt.

Zhang, et al., Kidney International, 56:549-558 (1999) identified genes up-regulated in 5/6 nephrectomized (subtotal renal ablation) mouse kidney by a PCR-based subtraction method. Ten known and nine novel genes were identified. The ultimate goal was to identify genes involved in glomerular hyperfiltration and hypertrophy.

Melia, et al., Endocrinol., 139:688-95 (1998) applied subtractive hybridization methods for the identification of androgen-regulated genes in mouse kidney. The treatment mice were dosed with dihydrotestosterone, an androgen. Kidney androgen-regulated protein gene was used as a positive control, as it is known to be up-regulated by DHT.

See also Holland, et al., Abstract 607, "Identification of Genes Possibly Involved in Nephropathy of Bovine Growth Hormone Transgenic Mice" (Endocrine Society Meeting, June 22, 2000) and Coschigano, et al., Abstract 333, "Identification of Genes Potentially Involved in Kidney Protection During Diabetes" (Endocrine Society Meeting, June 22, 2000).

The following differential hybridization articles may
also be of interest: Wada, et al., "Gene expression
profile in streptozotocin-induced diabetic mice kidneys

15

undergoing glomerulosclerosis", Kidney Int, 59:1363-73 (2001); Song, et al., "Cloning of a novel gene in the human kidney homologous to rat muncl3S: its potential role in diabetic nephropathy", Kidney Int., 53:1689-95 (1998); Page, et al., "Isolation of diabetes-associated kidney genes using differential display", Biochem. Biophys. Res. Comm., 232:49-53 (1997); Peradi, "Subtractive hybridization claims: An efficient technique to detect overexpressed mRNAs in diabetic nephropathy," Kidney Int. 53:926-31 (1998); Condorelli, EMBO J., 17:3858-66 (1998);

See also Nadler, S.T., Stoehr, J.P., Schueler, K.L., Tanimoto, G., Yandell, B.S., Attie, A.D. (2000) "The expression of adipogenic genes is decreased in obesity and diabetes mellitus", Proc Natl Acad Sci U S A 97:11371-11376; Lan H, Rabaglia ME, Stoehr JP, Nadler ST, Schueler KL, Zou F, Yandell BS, Attie AD. (2003) "Gene expression profiles of nondiabetic and diabetic obese mice suggest a role of hepatic lipogenic capacity in diabetes susceptibility", Diabetes 52:688-700.

See also WO00/66784 (differential hybridization screening for brown adipose tissue); PCT/US00/12366, filed May 5, 2000 (differential hybridization screening for liver).

Other Anti-Aging Studies

5

10

15

30.

15

0

5

For genes thought to have aging inhibitory activity, see generally International Longevity Center, Workshop Reports, "Longevity Genes: From Primitive Organisms to Humans," and "Is there an 'Anti-Aging' Medicine?".

Patents of possible interest include the following:

16

Lin, USP 6,303,768 (2001) ("Methuselah gene")

Lippman, USP 4,695,590 ("Method for retarding aging")

West, USP 6,368,789 (2002) ("Screening methods to identify inhibitors of telomerase activity")

10 Measurement of Biological Aging

Patents of possible interest include the following:

Kojima, USP 5,000,188 (1991) (an apparatus for measuring the physiological age of a subject).

15

20

25

Dimri, USP 5,795,728 (1998) ("Biomarkers of cell senescence")

Jia, USP 6,326,209 (2001) ("Measurement and quantification of 17 ketosteroid -sulfates as a biomarker of biological age")

Articles of interest include Kayo, et al., Proc. nat. Acad. Sci. (USA) 98:5093-98 (2001); Han, et al., Mch. Ageing Dev. 115:157-74 (2000); Dozmorov, et al., J. gerontol. A Biol. Sci. Med. Sci. 56:B72-B80 (2001); Dozmorov, et al., Id., 57: B99-B108 (2002); Miller, et al., Mol. Endocrinol., 16: 2657-66 (2002).

30

35

Apoptosis and CIDE-A

Apoptosis is a form of programmed cell death that occurs in an active and controlled manner that eliminates unwanted cells. Apoptotic cells undergo an orchestrated cascade of morphological changes such as membrane blebbing, nuclear shrinkage, chromatin condensation, and formation of

17

apoptotic bodies which then undergo phagocytosis by neighboring cells. One of the hallmarks of cellular apoptosis is the cleavage of chromosomal DNA into discrete oligonucleosomal size fragments. This orderly removal of unwanted cells minimizes the release of cellular components that may affect neighboring tissue. In contrast, membrane rupture and release of cellular components during necrosis often leads to tissue inflammation.

5

10

15

20

25

30

35

The process of apoptosis is highly conserved and involves the activation of the caspase cascade. Cohen, GM. (1997) Caspases: the executioners of apoptosis. Biochem. J. 326:1-16; Budihardjo, I., Oliver, H., Lutter, M., Luo, X., Wang, X. (1999) Biochemical pathways of caspase activation during apoptosis. Annnu. Rev. Cell. Dev. Biol.15:269-290; Jacobson, M.D., Weil, M., Raff, M.C. (1997) Programmed cell death in animal development. Cell 88:347-354. Caspases are a family of serine proteases that are synthesized as inactive proenzymes. Their activation by apoptotic signals such as CD95 (Fas) death receptor activation or tumor necrosis factor results in the cleavage of specific target proteins and execution of the apoptotic Apoptosis may occur by either an extrinsic pathway involving the activation of cell surface death receptors (DR) or by an intrinsic mitochondrial pathway. Yoon, J-H. Gores G.J. (2002) Death receptor-mediated apoptosis and the liver. J. Hepatology 37:400-410.

These pathways are not mutually exclusive and some cell types require the activation of both pathways for maximal apoptotic signaling. In type-I cells, death receptor activation leads to the recruitment and activation of caspases-8/10 and the rapid cleavage and activation of caspase-3 in a mitochondrial-independent manner.

Hepatocytes are members of the Type-II cells in which mitochondria are essential for DR-mediated apoptosis Scaffidi, C., Fulda, S., Srinivasan, A., Friesen, C., Li, F., Tomaselli, K.J., Debatin, K.M., Krammer, P.H., Peter,

18

M.E. (1998) Two CD95 (APO-1/Fas) signaling pathways. EMBO J. 17:1675-1687. In this pathway, the pro-apoptotic protein Bid is truncated activated caspases-8/10 and translocates to the mitochondria. Luo, X., Budihardjo, I., Zou, H., Slaughter, C., Wang, X. (1998) Bid, a Bcl2 interacting protein, mediates cytochrome c release from mitochondria in

protein, mediates cytochrome c release from mitochondria i response to activation of cell surface death receptors.

Cell 94:481-490; Li, H., Zhu, H., Xu, C.J., Yuan, J.

(1998) Cleavage of BID by caspase 8 mediates the

5

10

15

20

25.

30

35

mitochondrial damage in the Fas pathway of apoptosis. Cell 94:491-501. This translocation leads to mitochondrial cytochrome c release and eventual activation of caspases-3 and 7 via cleavage by activated caspase-9.

One of the substrates for activated caspase-3 is the DNA fragmentation factor (DFF). DFF is composed of a 45 kDa regulatory subunit (DFF45) and a 40 kDA catalytic subunit (DFF40). Liu, X., Zou, H., Slaughter, C., Wang, X. (1997) DFF, a heterodimeric protein that downstream of caspase-3 to trigger DNA fragmentation during apoptosis. Cell 89:175-184. DFF45 cleavage by activated caspase-3 results in its dissociation from DFF40 and allows the caspase-activated DNAse (CAD) activity of DFF40 to cleave chromosomal DNA into oligonucleosomal size fragments. Liu, X., Li, P., Widlak, P., Zou, H., Luo, X., Garrard, W.T., Wang, X. (1998) The 40-kDa subunit of DNA fragmentation factor induces DNA fragmentation and chromatin condensation during apoptosis. Proc. Natl. Acad. Sci. USA. 95:8461-8466; Halenbeck, R., MacDonald, H., Roulston, A., Chen, T.T., Conroy, L., Williams, L.T. (1998) CPAN, a human nuclease regulated by the caspase-sensitive inhibitor DFF45. Curr Biol. 8:537-540; Nagata, S. (2000) Apoptotic DNA fragmentation. Exp. Cell Res. 256:12-8.

Recently, a novel family of cell-death-inducing DFF45-like effectors (CIDEs) have been identified that includes CIDE-A, CIDE-B and CIDE-3/FSP2. Inohara, N., Koseki, T., Chen, S., Wu, X., Nunez, G. (1998) CIDE, a novel family of

19

cell death activators with homology to the 45 kDa subunit of the DNA fragmentation factor. EMBO J. 17:2526-2533; Danesch, U., Hoeck, W., Ringold, G.M. (1992) Cloning and transcriptional regulation of a novel adipocyte-specific gene, FSP27. CAAT-enhancer-binding protein (C/EBP) and C/EBP-like proteins interact with sequences required for differentiation-dependent expression. J. Biol. Chem. 267:7185-7193; Liang, L., Zhao, M., Xu, Z., Yokoyama, K.K., Li, T. (2003) Molecular cloning and characterization of CIDE-3, a novel member of the cell-death-inducing DNA-fragmentation-factor (DFF45)-like effector family. Biochem. J. 370:195-203.

5

10

15

20

The CIDEs contain an N-terminal domain that shares homology with the N-terminal region of DFF45 and may represent a regulatory region via protein interaction. See Inchara, supra; Lugovskoy, A.A., Zhou, P., Chou, J.J., McCarty, J.S., Li, P., Wagner, G. (1999) Solution structure of the CIDE-N domain of CIDE-B and a model for CIDE-N/CIDE-N interactions in the DNA fragmentation pathway of apoptosis. Cell 9:747-

- 755. The family members also share a C-terminal domain that is necessary and sufficient for inducing cell death and DNA fragmentation; see Inohara supra. The overexpression of CIDE-A induces cell death that can be inhibited by DFF45. However, CIDE-A-induced apoptosis in not inhibited by
- caspase-8 inhibitors thereby suggesting the presence of additional, caspase-independent, pathway(s) for the induction of apoptosis, see Inohara supra. Previous reports have indicated that human and mouse CIDE-A is expressed in several tissues such as brown adipose tissue (BAT) and heart and is localized to the mitochondria, Zhou, Z., Yon Toh, S., Chen, Z., Guo, K., Ng, C.P., Ponniah, S., Lin, S.C., Hong, W., Li, P. (2003) Cidea-deficient mice have lean phenotype and are resistant to obesity. Nat. Genet. 35:49-56. . In addition to the ability to induce apoptosis, CIDE-A can
- interact and inhibit UCP1 in BAT and may therefore play a role in regulating energy balance, see Zhou supra.

Previous reports have indicated that CIDE-A is not expressed in either adult human or mouse liver tissue, see Inohara supra, Zhou supra. We report here that CIDE-A is not only expressed in adult mouse liver tissue at older ages but is prematurely expressed in hyperinsulinemic and type-II diabetic mouse liver tissue. CIDE-A expression also correlates with liver steatosis in diet-induced obesity, hyperinsulinemia and type-II diabetes. These observations suggest an additional pathway of apoptotic cell death in NAFLD and that CIDE-A may play a role in this serious disease and potentially liver dysfunction associated with type-II diabetes.

5

10

15

20

25

30

35

SUMMARY OF THE INVENTION

Our attention recently has focused on the generation of liver mRNA expression profiles and the identification of genes involved in the aging process. We have therefore explored the genetic changes in the liver of C57Bl/6 mice that occur during the aging process, observing the gene expression patterns that occur at many different time points.

Gene chips have been used to identify mouse genes that are differentially expressed in mice, depending upon their age. We have utilized the Amersham product code: 300013 Codelink UniSet Mouse I Bioarray to determine the level of gene expression of approximately 10,000 mouse genes in the liver of mice with average ages of 35, 49, 77, 118, 133, 207, 403, 558 and 725 days.

In essence, complementary RNA derived from mice of different ages was screened for hybridization with oligonucleotide probes each specific to a particular mouse database DNA, as identified, by database accession number, by the gene manufacturer. Each database DNA in turn was also identified by the gene chip manufacturer as representative of a particular mouse gene cluster (Unique).

In most cases, this database DNA sequence was a full length genomic DNA or cDNA sequence, and are therefore either identical to, or encode the same protein as does, a natural full-length genomic DNA protein coding sequence. Those which don't at least present a partial sequence of a natural gene or its cDNA equivalent.

For the sake of simplicity, all of these mouse database DNA sequences, whether full-length or partial, and whether cDNA or genomic DNA, are referred to herein as "mouse genes". When only the genomic sequence is intended, we will refer specifically to "genomic DNA" or "qDNA".

The sequences in the protein databases are determined either by directly sequencing the protein or, more commonly, by sequencing a DNA, and then determining the translated

22

amino acid sequence in accordance with the Genetic Code.
All of the mouse sequences in the mouse polypeptide database are referred to herein as "mouse proteins" regardless of whether they are in fact full length sequences.

Mouse genes which were substantially differentially expressed (younger vs. older), as measured by different levels of hybridization of the respective cRNA samples with the particular probe corresponding to that mouse gene, were identified.

5

10

15

20

25

30

35

Favorable behavior is when expression decreases with age. Substantially favorable behavior is when the ratio of younger value to older value is at least two fold. Unfavorable behavior is when expression increases with age. Substantially unfavorable behavior is when the ratio of older value to younger value is at least two fold.

A mouse gene is considered to be "favorable" (more precisely, "wholly favorable") for the purpose of the Master Tables, especially subtable 1A, if, for at least one of the time comparisons set forth in the Examples, it exhibited substantially favorable behavior, and if, for all the other comparisons, it at least did not exhibit substantially unfavorable behavior. Note that the classification of a gene as favorable for purpose of the Master Table does not mean that it must have exhibited substantially favorable behavior for all of the comparisons set forth in the Examples.

A mouse gene is considered to be "unfavorable" (more precisely, "wholly unfavorable) for the purpose of the Master Tables, especially subtable 1B, if, for at least one of the time comparisons set forth in the Examples, it exhibited substantially unfavorable behavior, and if, for all the other comparisons, it at least did not exhibit substantially favorable behavior.

A mouse gene is considered to be "mixed" (in effect, both partially favorable and partially unfavorable) for the purpose of the Master Tables, especially subtable 1C, if for at least one of the time comparisons set forth in the

23

Examples it exhibited substantially favorable behavior and if for at least one of the other such comparisons it exhibited substantially unfavorable behavior.

The expression of a gene may first rise, then fall, with increasing age. Or it may first fall, and then rise. These are just the two simplest of several possible "mixed" expression patterns.

5

.0

0

5 :

0

Thus, we can subdivide the "favorables" into wholly and partially favorables. Likewise, we can subdivide the unfavorables into wholly and partially unfavorables. The genes/proteins with "mixed" expression patterns are, by definition, both partially favorable and partially unfavorable. In general, use of the wholly favorable or wholly unfavorable genes/proteins is preferred to use of the partially favorable or partially unfavorable ones.

It is evident from the foregoing that mixed genes/proteins are those exhibiting a combination of favorable and unfavorable behavior. A mixed gene/protein can be used as would a favorable gene/protein if its favorable behavior outweighs the unfavorable. It can be used as would an unfavorable gene/protein if its unfavorable behavior outweighs the favorable. Preferably, they are used in conjunction with other agents that affect their balance of favorable and unfavorable behavior. Use of mixed genes/proteins is, in general, less desirable than use of purely favorable or purely unfavorable genes/proteins.

It will be appreciated that the comparisons set forth in the Examples are not exhaustive and that it is possible that a mouse gene which, on the basis of those comparisons, was classified as a "favorable" gene in the Master Table may turn out, if additional time points are considered, to sometimes exhibit substantially unfavorable behavior.

Nonetheless, such a gene will still be considered a "favorable" gene for the purpose of the Master Table and the claims referring to the Master Table. Likewise, a gene which, on the basis of those comparisons, was classified as

24

an "unfavorable" gene in the Master Table may prove, under more detailed examination, to sometimes exhibit substantially favorable behavior. Nonetheless, it will retain "unfavorable" classification for the purpose of the Master Table and the claims referring thereto.

5

LO

L5

30

:5

0

.5

The "favorable", "unfavorable" and "mixed" mouse proteins are thus those listed in the Master Table as encoded by the listed "favorable", "unfavorable" and "mixed" mouse genes, respectively, or which otherwise correspond to those mouse genes.

Related human genes (database DNAs) and proteins were identified by searching a database comprising human DNAs or proteins for sequences corresponding to (i.e., homologous to, i.e., which could be aligned in a statistically significant manner to) the mouse gene or protein. The "favorable", "unfavorable" and "mixed" human genes and proteins are those which correspond to the listed "favorable", "unfavorable" and "mixed" mouse genes and proteins, respectively. More than one human protein may be identified as corresponding to a particular mouse chip probe and to a particular mouse gene.

Note that the terms "human genes" and "human proteins" are used in a manner analogous to that already discussed in the case of "mouse genes" and "mouse proteins", e.g., the "genes" include both gDNA and cDNA, and both full and partial sequences.

As used herein, the term "corresponding" does not mean identical, but rather implies the existence of a statistically significant sequence similarity, such as one sufficient to qualify the human protein or gene as a homologous protein or DNA as defined below. The greater the degree of relationship as thus defined (i.e., by the statistical significance of each alignment used to connect the mouse chip DNA, and the corresponding mouse gene/cDNA, to the human protein or gene, measured by an E value), the more close the correspondence. The connection may be direct

25

(mouse gene to human protein) or indirect (e.g., mouse gene to human gene, human gene to human protein). By "mouse gene", we mean the mouse gene from which the gene chip DNA in question was derived.

In general, the human genes/proteins which most closely correspond, directly or indirectly, to the mouse genes are preferred, such as the one(s) with the highest, top two highest, top three highest, top four highest, top five highest, and top ten highest E values for the final alignment in the connection process. The human genes/proteins deemed to correspond to our mouse genes are identified in the Master Tables.

10

15

20

25

30

35

Note that it is possible to identify homologous fulllength human genes and proteins, if they are present in the database, even if the query mouse DNA or protein sequence is not a full-length sequence.

If there is no homologous full-length human gene or protein in the database, but there is a partial one, the latter may nonetheless be useful. For example, a partial protein may still have biological activity, and a molecule which binds the partial protein may also bind the full-length protein so as to antagonize a biological activity of the full-length protein. Likewise, a partial human gene may encode a partial protein which has biological activity, or the gene may be be useful in the design of a hybridization probe or in the design of a therapeutic antisense DNA.

The partial genes and protein sequences may of course also be used in the design of probes intended to identify the full length gene or protein sequence.

Agents which bind the "favorable" and "unfavorable" nucleic acids (e.g., the agent is a substantially complementary nucleic acid hybridization probe), or the corresponding proteins (e.g., an antibody vs. the protein) may be used to estimate the biological age of a human

26

subject, or to predict the rate of biological aging in a human subject (i.e, to evaluate whether a human subject is at increased or decreased risk for faster-than-normal biological aging.) A subject with one or more elevated "unfavorable" and/or one or more depressed "favorable" genes/proteins is at increased risk, and one with one or more elevated "favorable" and/or one or more depressed "unfavorable" genes/proteins is at decreased risk.

The assay may be used as a preliminary screening assay to select subjects for further analysis, or as a formal diagnostic assay.

The identification of the related genes and proteins may also be useful in protecting humans against faster-thannormal or even normal aging (hereinafter, "the disorders").
They may be used to reduce a rate of biological aging in the subject, and/or delay the time of onset, or reduce the severity, of an undesirable age-related phenotype in said subject, and/or protect against an age-related disease.

<u>}</u>0

:5

0

5

5

LO

15

Thus, Applicants contemplate:

- (1) use of the "favorable" mouse DNAs (or fragments thereof) of the Master Tables (below) to isolate or identify related human DNAs;
- (2) use of human DNAs, related to favorable mouse DNAs, to express the corresponding human proteins;
- (3) use of the corresponding human proteins (and mouse proteins, if biologically active in humans), to protect against the disorder(s);
- (4) use of the corresponding mouse or human proteins, or nucleic acid probes derived from the mouse or human genes, in diagnostic agents, in assays to measure or predict biological aging or the rate thereof; and
- (5) use of the corresponding human or mouse genes therapeutically in gene therapy, to protect against the disorder(s).

27

Moreover Applicants contemplate:

5

10

15

(1) use of the "unfavorable" mouse DNAs (or fragments thereof) of the Master Tables to isolate or identify related human DNAs;

- (2) use of the complement to the "unfavorable" mouse DNAs or related human DNAs, as antisense molecules to inhibit expression of the related human DNAs;
- (3) use of the mouse or human DNAs to express the corresponding mouse or human proteins;
- (4) use of the corresponding mouse or human proteins, in diagnostic agents, to measure biological aging or the rate thereof;
- (5) use of the corresponding mouse or human proteins in assays to determine whether a substance binds to (and hence may neutralize) the protein; and
- (6) use of the neutralizing substance to protect against the disorder(s).

Thus, DNAs of interest include those which specifically 20 hybridize to the aforementioned mouse or human genes, and are thus of interest as hybridization assay reagents or for antisense therapy. They also include synthetic DNA sequences which encode the same polypeptide as is encoded by the database DNA, and thus are useful for producing the polypeptide in cell culture or in situ (i.e., gene therapy). 25 Moreover, they include DNA sequences which encode polypeptides which are substantially structurally identical or conservatively identical in amino acid sequence to the mouse and human proteins identified in the Master Table 1, subtables 1A or 1C, and DNA sequences which encode human 30 proteins which are members of human protein classes set forth in master table 2, subtables 2A or 2C. Finally, they include DNA sequences which peptide (including antibody) antagonists of the proteins of Master Table 1, subtables 1B or 1C, or of human proteins which are members of human 35 protein classes set forth in master table 2, subtables 2B or

28

2C.

5

10

15

Related human DNAs also may be identified by screening human cDNA or genomic DNA libraries using the mouse gene of the Master Table, or a fragment thereof, as a probe.

If the mouse gene of Master Table 1 is not full-length, and there is no closely corresponding full-length mouse gene in the sequence databank, then the mouse DNA may first be used as a hybridization probe to screen a mouse cDNA library to isolate the corresponding full-length sequence.

Alternatively, the mouse DNA may be used as a probe to screen a mouse genomic DNA library.

The human protein cell death activator CIDE-A is of particular interest because of its highly dramatic change in liver expression with age.

20

25

The agents of the present invention may be used alone or in conjunction with each other and/or known anti-aging or anti-age-related disease agents. It is of particular interest to use the agents of the present invention in conjunction with an agent disclosed in one of the related applications cited above.

BRIEF DESCRIPTION OF THE DRAWINGS

- Fig. 1 CIDE-A Expression is elevated in older normal mice. CIDE-A expression is plotted for normal C57BI/6J mouse ages 35, 49, 56, 77, 133, 207, 403, 558 and 725 days. Expression is low for the first few data points, then rises sharply at 403 days, and again at 558 days. There is a drop off at 725 days, but expression remains above the 403 day evel.
- 3. 2 CIDE-A Expression is elevated at an earlier age in abetic mice. In diabetic mice, the CIDE-A expression at .33 days is more than double that at 77 days, while in normal mice, the increase over the same interval is slight.
- Fig. 3. Steatosis in liver of high-fat diet fed mice. Mice were weaned directly onto either a normal diet or a high-fat diet and maintained on the respective diets for up to 26 weeks. The mice were sacrificed and liver tissue isolated. Percent liver white space was determined.

30

DETAILED DESCRIPTION OF THE PREFERRED EMBODIMENTS OF THE INVENTION

Full-Length vs. Partial Length Genes/Proteins

A "full length" gene is here defined as a (1) a naturally occurring DNA sequence which begins with an initiation codon (almost always the Met codon, ATG), and ends with a stop codon in phase with said initiation codon (if introns, if any, are ignored), and thereby encodes a naturally occurring polypeptide with biological activity, or a naturally occurring precursor thereof, or (2) a synthetic DNA sequence which encodes the same polypeptide as that which is encoded by (1). The gene may, but need not, include introns.

A "full-length" protein is here defined as a naturally occurring protein encoded by a full-length gene, or a protein derived naturally by post-translational modification of such a protein. Thus, it includes mature proteins, proproteins, preproteins and preproproteins. It also includes substitution and extension mutants of such naturally occurring proteins.

25 Subjects

30

35

5

10

For mice, infancy is defined as the period 0 to 21 days after birth. Sexual maturity is reached, on average, at 42 days after birth. The average lifespan is 832 days.

In humans, infancy is defined as the period between birth and two years of age. Sexual maturity in males can occur between 9 and 14 years of age while the average age at first menstrual period for females 15-44 years old is 12.6 years. The average human lifespan is 73 years for males and 79 years for females. The maximum verified human lifespan was 122 years, five months and 14 days.

31 Chronological and Biological Aging

5

10

15

20

25

10

5

'Aging" is a process of gradual and spontaneous change, resulting in maturation through childhood, puberty, and young adulthood and then primarily a decline in function through middle and late age. Aging thus has both the positive component of development/maturation and the negative component of decline.

"Senescence" refers strictly to the undesirable changes that occur as a result of post-maturation aging. Some of the changes which occur in post-maturation aging are not deleterious to health (e.g., gray hair, baldness), and some may even be desirable (e.g., increased wisdom and experience). In contrast, the memory impairment that occurs with age is considered senescence. However, we will hereafter use "aging" per se to refer to "senescence", and use "maturation" to refer to pre-maturation development.

There is increased mortality with age after maturation. There is also a progressive decrease in physiological capacity with age, but the rate of physiological decline varies from organ to organ and from individual to individual. The physiological decline results in a reduced ability to respond adaptively to environmental stimuli, and increased susceptibility and vulnerability to disease.

"Aging is the accumulation of diverse adverse changes that increase the risk of death. These changes can be attributed to development, genetic defects, the environment, disease, and the inborn aging process. The chance of death at a given age serves as a measure of the number of accumulated changes, that is, of physiologic age, and the rate of change of this measure, as the rate of aging."

Harman, Ann. N.Y. Acad. Sci. 854:1-7 (1998).

Preferably, the agents of the present invention inhibit aging for at least a subpopulation of mature (post-puberty) adult subjects.

The term "healthy aging" (sometimes called "successful aging") refers to post-maturation changes in the body that

32

occur with increasing age even in the absence of an overt disease. However, increased age is a risk factor for many diseases ("age-related diseases"), and hence "total aging" includes both the basal effects of healthy aging and the effects of any age-related disease. (Most literature uses the term "normal aging" as a synonym for "healthy aging", but a minority use it to refer to "total aging". To minimize confusion, we will try to avoid the term "normal aging", but if we use it, it is as a synonym for "healthy aging".) Some scientists have suggested that normal aging changes should be defined as those which are universal, degenerative, progressive and intrinsic.

-10

. 5

: 0

Preferably, the agents of the present invention inhibit healthy aging for at least a subpopulation of mature (post-puberty) adult subjects.

In both aging and senescence, many physiologic functions decline, but normal decline is not usually considered the same as disease. The distinction between normal decline and disease is often but not always clear and may be due only to statistical distribution. Glucose intolerance is considered consistent with healthy aging, but diabetes is considered a disease, although a very common one. Cognitive decline is nearly universal with advanced age and is considered healthy aging; however, cognitive decline consistent with dementia, although common in late life, is considered a disease (as in the case of Alzheimer's, a conclusion supported by analysis of brain tissue at autopsy). A decline in maximal heart rate is typical of healthy aging. In contrast, coronary heart disease is an age-related disease. A decline in bone density is considered healthy aging, but when it drops to 2.5 SD below the young adult mean, it is called osteoporosis. Generally speaking, the changes typical of healthy aging are gradual, while those typical of a disorder can be rapid.

The term average (median) "lifespan" is the chronological age to which 50% of a given population

33

survive. The maximum lifespan potential is the maximum age achievable by a member of the population. As a practical matter, it is estimated as the age reached by the longest lived member (or former member) of the population. The (average) life expectancy is the number of remaining years that an individual of a given age can expect to live, based on the average remaining lifespans of a group of matched individuals.

The most widely accepted method of measuring the rate of aging is by reference to the average or the maximum lifespan. If a drug treatment achieves a statistically significant improvement in average or maximum lifespan in the treatment group over the control group, then it is inferred that the rate of aging was retarded in the treatment group. Similarly, one can compare long-term survival between the two groups.

10

15

30

35

10

15

Preferably, the agents of the present invention have the effect of increasing the average lifespan and/or the maximum lifespan for at least a subpopulation of mature (post-puberty) adult subjects. This subpopulation may be defined by sex and/or age. If defined in part by age, then it may be defined by a minimum age (e.g., at least 30, at least 40, at least 50, at least 55, at least 60, at least 65, at least 70, at least 75, at least 80, at least 90, etc.) or by a maximum age (not more than 40, not more than 50, not more than 55, not more than 60, not more than 65, not more than 70, not more than 75, not more than 80, not more than 90, not more than 100, etc.), or by a rational combination of a minimum age and a maximum age so as to define a preferred close-ended age range, e.g., 55-75.

The subpopulation may additionally be defined by race, e.g., caucasian, negroid or oriental, and/or by ethnic group, and/or by place of residence (e.g., North America, Europe).

The subpopulation may additionally be defined by nonage risk factors for age-associated diseases, e.g., by blood

34

pressure, body mass index, etc.

Preferably, the subpopulation in which an agent of the present invention is reasonably expected to be effective is large, e.g., in the United States, preferably at least 100,000 individuals, more preferably at least 1,000,000 individuals, still more preferably at least 10,000,000, even more preferably at least 20,000,000, most preferably at least 40,000,000.

By way of comparison, according to the 2000 U.S. Census, the U.S. population, by age, was

Age	Pop (mil)
15-19	20.2
20-24	19.0
25-29	19.4
30-34	20.5
35-39	22.7
40-44	22.4
45-49	20.1
50-54	17.6
55-59	13.5
60-64	10.8
65-69	9.5
70-74	8.9
75-79	7.4
80-84	4.9
85+	4.2

LO

For any given chronological age, statisticians can define the probability of living to a particular later age. These expectancies can be calculated for the entire age cohort, or broken down by sex, race, country of residence, etc. Individuals who live longer than expected can be said, after the fact, to have biologically aged more slowly

35

than their peers. One definition of biological age is that it is a measure of one's position in one's life span, i.e., biological age = position in own life span (as fraction in range 0..1) X average life span for species. This simple definition carries with it the implicit assumption that the rate of biological aging is constant. It also has the practical problem of determining one's own life span before death. We will present a more practical definition shortly.

5

.0

5

0

5 '

0

5

The problem with lifespan studies is that they are extremely time-consuming. A maximum lifespan study in mice can take 4-5 years. A maximum lifespan study in dogs or cats would take 15-20 years, in monkeys, 30-40 years, and in humans, over 100 years. Even if the human study group were of sexagenarians, it would take 40-60 years to complete the study.

Hence, scientists have sought to identify biological markers (biomarkers) of biological aging, that is, characteristics that can be measured while the subjects are still alive, which correlate to lifespan. These biological markers can be used to calculate a "biological age" (syn. "Physiological age"); it is the chronological age at which an average member of the population (or relevant subpopulation) would have the same value of a biomarker of biological aging (or the same value of a composite measure of biomarkers of biological aging) as does the subject. This is the definition that will be used in this disclosure, unless otherwise stated.

The effect of aging varies from system to system, organ to organ, etc. For example, between ages 30 and 70 years, nerve conduction velocity decreases by only about 10%, but renal function decreases on average by nearly 40%. Thus, there isn't just one biological age for a subject. By a suitable choice of biomarker, one may obtain a whole organism, or a system-, organ- or tissue-specific measure of biological aging, e.g., one can say that a person has the nervous system of a 30 year old but the renal system of a 60

36

year old. Biomarkers may measure changes at the molecular, cellular, tissue, organ, system or whole organism levels.

Generally speaking, in the absence of some form of intervention (drugs, diet, exercise, etc.), biological ages will increase with time. The agents of the present invention preferably reduce the time rate of change of a biological age of the subject. The term "a biological age" could refer to the overall biological age of the subject, to the biological age of a particular system, organ or tissue of that subject, or to some combination of the foregoing. More preferably, the agents of the present cannot only reduce the rate of increase of a biological age of the subject, but can actually reduce a biological age of the subject.

5

10

15

30

25

10

5

A simple biologic marker (biomarker) is a single biochemical, cellular, structural or functional indicator of an event in a biologic system or sample. A composite biomarker is a mathematical combination of two or more simple biomarkers. (Chronological age may be one of the components of a composite biomarker.)

A plausible biomarker of biological age would be a biomarker which shows a cross-sectional and/or longitudinal correlation with chronological age. Nakamura suggests that it is desirable that a biomarker show (a) significant cross-sectional correlation with chronological cage, (b) significant longitudinal change in the same direction as the cross-sectional correlation, (c) significant stability of individual differences, and (d) rate of age-related change proportional to differences in life span among related species. Cp. Nakamura, Exp Gerontol. 29(2):151-77 (1994), using desiderata (a)-(c). A superior biomarker of biological age would be a better predictor of lifespan than is chronological age (preferably for a chronological age at which 90% of the population is still alive).

The biomarker preferably also satisfies one or more of

37

the following desiderata: a statistically significant agerelated change is apparent in humans after a period of at most a few years; not affected dramatically by physical conditioning (e.g., exercise), diet, and drug therapy (unless it is possible to discount these confounding influences, e.g., by reference to a second marker which measures them); can be tested repeatedly without harming the subject; works in lab animals as well as humans; simple and inexpensive to use; does not alter the result of subsequent tests for other biomarkers if it is to be used in conjunction with them; monitors a basic process that underlies the aging process, not the effects of disease.

5

10

L5

30

:5

ດັ

5

Preferably, if the biomarker works in lab animals, there is a statistically significant difference in the value of the biomarker between groups of food-restricted and normally-fed animals. It has been shown in some mammalian species that dietary restriction without malnutrition (e.g., caloric decrease of up to 40% from ad libitum feeding) increases lifespan.

A biomarker of aging may be used to predict, instead of lifespan, the "Healthy Active Life Expectancy" (HALE) or the "Quality Adjusted Life Years" (QALY), or a similar measure which takes into account the quality of life before death as well as the time of death itself. For HALE, see Jagger, in Outcomes Assessment for Healthcare in Elderly People, 67-76 (Farrand Press: 1997). For QALY, see Rosser RM. A health index and output measure, in Stewart SR and Rosser RM (eds) Quality of Life: Assessment and Application. Lancaster: MTP, 1988.

A biomarker of aging may be used to predict, instead of lifespan, the timing and/or severity of a change in one or more age-related phenotypes as described below.

A biomarker of aging may be used to estimate, rather than overall biological age for a subject, a biological age for a specific body system or organ. The determination of the biological age of the liver, and the inhibition of

38

biological aging of the liver, are of particular interest.

Body systems include the nervous system (including the brain, the sensory organs, and the sense receptors of the skin), the cardiovascular system (includes the heart, the red blood cells and the reticuloendothelial system), the respiratory system, the gastrointestinal system, the endocrine system (pituitary, thyroid, parathyroid and adrenal glands, gonads, pancreas, and parganglia), the musculoskeletal system, the urinary system (kidneys, bladder, ureters, urethra), the reproductive system and the immune system (bone marrow, thymus, lymph nodes, spleen, lymphoid tissue, white blood cells, and immunoglobulins). biomarker may be useful in estimating the biological age of a system because the biomarker is a chemical produced by that system, because it is a chemical whose activity is primarily exerted within that system, because it is indicative of the morphological character or functional activity of that system, etc. A given biomarker may be thus associated with more than one system. In a like manner, a biomarker may be associated with the biological age, and hence the state, of a particular organ or tissue.

15

:0

0

5

The prediction of lifespan, or of duration of system or organ function at or above a particular desired level, may require knowledge of the value of at least one biomarker of aging at two or more times, adequately spaced, rather than of the value at a single time. See McClearn, Biomarkers of Age and Aging, Exp. Gerontol., 32:87-94 (1997).

The levels (or changes in levels) of the human proteins identified in this specification, and their corresponding mRNAs, may be used as simple biomarkers (direct or inverse) of biological aging. They may be used in conjunction with each other, or other simple biomarkers, in a composite biomarker.

Once several plausible simple biomarkers have been

39

identified, a composite biomarker may be obtained by standard mathematical techniques, such as multiple regression, principal component analysis, cluster analysis, neural net analysis, and so forth. As a preliminary to such analysis, the values may be standardized, e.g., by converting the raw scores into z-scores based on the distributions for each simple biomarker.

For example, principal component analysis can be used to analyze the variation of lifespan with different observables, and the factor score coefficients from the first principal component can be used to derive an equation for estimating a biological age score. Nakamura, Exp Gerontol. 29(2):151-77 (1994). This approach was used to obtain the following BAS (for healthy Japanese women aged 28-80): BAS=-4.37 -0.998FEV_{1.0} +0.022SBP +0.133MCH +0.018GLU -1.505 A/G RATIO, where FEV, o is the forced expiratory volume in 1 sec. (Liters), SBP is the systolic blood pressure (mm Hg), MCH is the mean corpuscular hemoglobin (pq), GLU is glucose (mq/dl), and A/G RATIO is the ratio of albumin to globulin. The relative importance of these five biomarkers was 33.7%, 25.1%, 17.1%, 14.8% and 8.9%, respectively. Ueno, et al., "Biomarkers of Aging in Women and the Rate of Longitudinal Changes," J. Physiol. Anthropol. 22(1): 37-46 (Jan. 2003).

It should be noted that particularly when evaluating the overall biological age of the subject, it is not necessarily most desirable to weight all systems or all organs equally. One may find it more desirable to give greater weight to the system or organ with the highest biological age in calculating the overall biological age, because it is presumably more likely to deteriorate or fail, resulting in death. Appropriate statistical analysis can be used to find the weighting scheme resulting in the best prediction of lifespan.

5

10

15

20

25

30

40

simple biomarkers is used to measure human aging:

SENSORY

- 1. Highest audible pitch (kHz)
- 2. Visual accommodation (diopters)
 - 3. Vibrotactile sensitivity (dB)

MOTOR

- 4. Muscle Movement time (sec)
- 10 5. Muscle Movement time with decision (sec)
 - 6. Alternate button tapping time (sec)

COGNITIVE

- 7. Memory, length of sequence
- 15 8. Auditory reaction time (sec)
 - 9. Visual reaction time (sec)
 - 10. Visual Reaction time with decision (sec)

PULMONARY

20

25

3 0

35

- 11. Forced vital capacity (liters)
 - 12. Forced expiratory Volume- 1 sec (liters)

See Hochschild, R., Journal of Gerontology [Biological Science] 45(6):B187-214; 1990).

According to a website discussing the H-SCAN test, "Biomarkers of aging are characteristics of an organism that correlate in large groups with chronological age and correlate in large groups with chronological age and correlations are biomarkers of aging that also correlate with the quality of life in later life in the sense that they involve functions that are crucial to carrying out the activities of daily living... A single biomarker of aging is limited by the fact that it measures only one isolated characteristic and is hardly representative of the diversity of functional and structural concomitants of aging... Biological age, in

41

contrast to chronological age, is an individual's hypothetical age calculated from scores obtained on a battery of tests of biomarkers of aging. As a first step in the calculation, the age of which each biomarker score is typical is determined by comparison with scores obtained by a large representative group of persons (or organisms) spanning a range of ages. Then one of a variety of averaging techniques is employed (optionally with standardization steps) to obtain a single index of age, as described in detail by Hochschild. This index varies with, and therefore must be expressed with reference to, the measured biomarkers and the mathematical method of combining scores."

http://www.longevityinstituteone.com/

5

10

15

20

25

30

35

Abbo, USP 6,547,729 teaches determining the biological age (he calls it "performance age") of a subject by (1) for a sample population, determining a regression curve relating some set of observed values for an "indicator" of the functionality of a bodily system to the chronological age of the observed individuals, (2) solving the regression equation to obtain a predicted performance age, given the value of the indicator for the subject. The regression can be based on more than one indicator, i.e., it can be a multiple regression. The sample population can be defined by sex, age range, ethnic composition, and geographic The bodily system may be a molecular, cellular, location. tissue or organ system. The following indicators are suggested by Abbo: nervous system (memory tests, reaction time, serial key tapping, digit recall test, letter fluency, category fluency, nerve conduction velocity), arteries (pulse wave velocity; ankle-brachial index), skeletal system (bone mineral density); lungs (forced vital capacity), heart (ejection fraction; length of time completed on a treadmill stress test), kidneys (creatinine clearance), proteins (glycosylation of hemoglobin), endocrine glands (load level of bioactive testosterone; level of dehydroepiandrosterone

42

sulfate, ratio of urinary 17-ketosteroids/17-hydroxycorticosteroids; growth hormone; IGF-1).

Preferably, the agents of the invention have a favorable effect on the value of at least one simple biomarker of biological aging, such as any of the plausible biomarkers mentioned anywhere in this specification, other than the level of one of the proteins of the present invention. More preferably, they have a favorable effect on the value of at least two such simple biomarkers of biological aging. Even more preferably, at least one such pair is of markers which are substantially non-correlated ($R^2 < 0.5$).

Desirably, if more than one simple biomarker is favorably affected, the biomarkers in question reflect different levels of organization, and/or different body components at the same level of organization. For example, a visual reaction time with decision test is on the whole organism level, while a measurement of telomere length is on the cellular level.

A biomarker may, but need not, be an indicator related to one of the postulated causes or contributing factors of aging. It may, but need not, be an indicator of the acute health of a particular body system or organ.

A biomarker may measure behavior, cognitive or sensory function, or motor activity, or some combination thereof. It may measure the level of a type of cell (e.g., a T cell subset, such as CD4, CD4 memory, CD4 naive, and CD4 cells expressing P-glycoprotein) or of a particular molecule (e.g., growth hormone, IGF-1, insulin, DHEAS, an elongation factor, melatonin) or family of structurally or functionally related molecules in a particular body fluid (especially blood) or tissue. For example, lower serum IGF-1 levels are correlated with increasing age, and IGF-1 is produced by

43

many different tissues. On the other hand, growth hormone is produced by the pituitary gland.

A biomarker may measure an indicator of stress (particularly oxidative stress) and resistance thereto. It has been theorized that free radicals damage biomolecules, leading to aging.

A biomarker may measure protein glycation or other protein modification (e.g., collagen crosslinking). It has been theorized that such modifications contribute to aging.

The biomarker may measure changes in the lengths of telomeres or in the rate of cell division. It has been theorized that telomere shortening beyond a critical length leads the cell to stop proliferating. Average telomere length therefore provides a biomarker as to how may divisions the cell as previously undergone and how many divisions the cell can undergo in the future.

Suggested biomarkers have also included resting heart rate, resting blood pressure, exercise heart rate, percent body fat, flexibility, grip strength, push strength, abdominal strength, body temperature, and skin temperature.

The present invention does not require that all of the biomarkers identified above be validated as indicative of biological age, or that they be equally useful as measures of biological age.

5

0

5

5

.0

.5

:0

There is an overlap between biomarkers of aging and indicators of functional status. An indicator of functional status is an indicator that defines a functional ability (e.g., physiological, cognitive or physical function). An indicator of functional status may also be related to the increase in morbidity and mortality with chronological age. Such indicators preferably predict physiological, cognitive and physical function in an age-coherent way, and do so better than chronological age. Preferably, they can predict the years of remaining functionality, and the trajectory toward organ-specific illness in the individual. Also, they

44

are preferably minimally invasive.

Suggested indicators include anthropometric data (body mass index, body composition, bone density, etc.), functional challenge tests (glucose tolerance, forced vital capacity), physiological tests (cholesterol/HDL, glycosylated hemoglobin, homocysteine, etc.) and proteomic tests.

A number of mouse models for human aging exist. See Troen, supra, Table 3. The drugs identified by the present invention may be further screened in one or more of these models.

Age-Related Phenotype

An age-related phenotype is an observable change which occurs with age. An age-related phenotype may, but need not, also be a biomarker of biological aging.

Preferably, the agent of the present invention favorably affects at least one age-related phenotype. More preferably, it favorably affects at least two age-related phenotypes, more preferably phenotypes of at least two different body systems.

The age-related phenotype may be a system level phenotype, such as a measure of the condition of the nervous system, respiratory system, immune system, circulatory system, endocrine system, reproductive system, gastrointestinal system, or musculoskeletal system.

The age-related phenotype may be an organ level phenotype, such as a measure of the condition of the brain, eyes, ears, lungs, spleen, heart, pancreas, liver, ovaries, testicles, thyroid, prostate, stomach, intestines, or kidney.

45

The age-related phenotype may be a tissue level phenotype, such as a measure of the condition of the muscle, skin, connective tissue, nerves, or bones.

- The age-related phenotype may be a cellular level phenotype, such as a measure of the condition of the cell wall, mitochondria or chromosomes.
- The age-related phenotype may be a molecular level

 phenotype, such as a measure of the condition of nucleic
 acids, lipids, proteins, oxidants, and anti-oxidants.

The age-related phenotype may be manifested in a biological fluid, such as blood, urine, saliva, lymphatic fluid or cerebrospinal fluid. The biochemical composition of these fluid may be an overall, system level, organ level, tissue level, etc. phenotype, depending on the specific biochemical and fluid involved.

PHYSIOLOGICAL AGING OF THE HUMAN BODY BY SYSTEMS

SKIN, HAIR,	Loss of subcutaneous fat, Thinning of skin,
NAILS	Decreased collagen, Nails brittle and flake,
	Mucous membranes drier, Less sweat glands,
	Temperature regulation difficult, Hair
	pigment decreases, Hair thins. Eyelids baggy
	and wrinkled.

	46
EYES AND	Eyes deeper in sockets; Conjunctiva thinner
VISION	and yellow; Quantity of tears decreases; Iris
	fades; Pupils smaller, let in less light;
	Night and depth vision less; "Floaters" can
	appear
	Lens enlarges; Lens becomes less transparent,
	can actually become clouded, results in
	cataracts; Accommodation decreases, results
	in presbyopia; Impaired color vision, also -
	especially greens and blues because cones
	degenerate; Predisposed to glaucoma
	(Increased pressure in eye, decreased
	absorption of intraocular fluid; can result
	in blindness);
ł	Macular degeneration becoming more frequent
	(This is the patch of retina where lens
	focuses light, Ultimately results in
	blindness)
EARS AND	Irreversible, sensorineural loss
HEARING LOSS	(presbycusis) with age (Men more affected
	than women, Loss occurs in higher range of
	sound, By 60 years, most adults have trouble
	hearing above 4000Hz, Normal speech
	500-2000Hz)
RESPIRATORY	Lungs become more rigid, Pulmonary function
SYSTEM	decreases, Number and size of alveoli
	decreases, Vital capacity declines, Reduction
	in respiratory fluid, Bony changes in chest
	cavity
CARDIOVASCUL	Heart smaller and less elastic with age, By
AR SYSTEM	age 70 cardiac output reduced 70%, Heart
	valves become sclerotic, Heart muscle more
	irritable, More arrhythmias, Arteries more
	rigid, Veins dilate

	4 /
GASTROINTEST	Reduced GI secretions, Reduced GI motility,
INAL SYSTEM	Decreased weight of liver,
	Reduced regenerative capacity of liver, Liver
	metabolizes less efficiently
RENAL SYSTEM	After 40 renal function decreases, By 90 lose
	50% of function, Filtration and reabsorption
	reduced, Size and number of nephrons
	decrease,
	Bladder muscles weaken, Less able to clear
	drugs from system, Smaller kidneys and
	bladder
REPRODUCTIVE	Reduced testosterone level, Testes atrophy
SYSTEM	and soften, Decrease in sperm production,
(MALE)	Seminal fluid decreases and more viscous,
	Erections take more time, Refractory period
	after ejaculation may lengthen to days
REPRODUCTIVE	Declining estrogen and progesterone levels,
SYSTEM	Ovulation ceases, Introitus constricts and
(FEMALE)	loses elasticity, Vagina atrophies - shorter
	and drier,
-	Uterus shrinks, Breasts pendulous and lose
	elasticity
NEUROLOGICAL	Neurons of central and peripheral nervous
SYSTEM	system degenerate, Nerve transmission slows,
	Hypothalamus less effective in regulating
	body temperature, Reduced REM sleep,
	decreased deep sleep, After age 50, lose 1%
	of neurons each year
MUSCULOSCELE	Adipose tissue increases with age, Lean body
TAL SYSTEM	mass decreases, Bone mineral content
	diminished, Decrease in height from narrow
	vertebral spaces, Less resilient connective
	tissue, Synovial fluid more viscous,
	May have exaggerated curvature of spine

5

10

48

IMMUNE	Decline in immune function, Trouble
SYSTEM	differentiating between self and non-self -
	more auto-immune problems, Decreases antibody
	response,
	Fatty marrow replaced red marrow, Vitamin B12
	absorption might decrease - decreased
	hemoglobin and hematocrit
ENDOCRINE	Decreased ability to tolerate stress - best
SYSTEM	seen in glucose metabolism,
	Estrogen levels decrease in women, Other
	hormonal decreases include testosterone,
	aldosterone, cortisol, progesterone

Adapted from http://www.texashste.com/html/ger-pap1.ppt

5

10

20

25

The aging human liver appears to preserve its morphology and function relatively well. The liver appears to progressively decrease in both mass and volume. It also appears browner (a condition called "brown atrophy"), as a result of accumulation of lipofuscin (ceroid) within hepatocytes. Increases occur in the number of macrohepatocytes, and in polyploidy, especially around the terminal hepatic veins. The number of mitochondria declines, and both the rough and smooth endoplasmic recticulum diminish. The number of lysozymes increase.

The liver is the premiere metabolic organ of the body. With regard to metabolism, hepatic glycerides and cholesterol levels increase with age, at least up to age 90. On the other hand, phospholipids, aminotransferases, and serum bilirubin appear to remain normal. There are contradictory reports as to the effect of aging on albumin, serum gamma-glutamyltransferase, and hepatic alkaline phosphatase. It is worth noting that it has been shown that the content of cytochrome oxidase exhibits a progressive decline which correlates with age-associated decline in mtRNA synthesis in brain, liver, heart, lungs and skeletal

49

muscle.

See generally Anaantharaju, Feller and Chedid, "Aging Liver: A Review," Gerontology, 48: 343-53 (2002).

5 Quality of Life

10

25

30

15

Clinicians are interested, not only in simple prolongation of lifespan, but also in maintenance of a high quality of life (QOL) over as much as possible of that lifespan. QOL can be defined subjectively in terms of the subject's satisfaction with life, or objectively in terms of the subject's physical and mental ability (but not necessarily willingness) to engage in "valued activities", such as those which are pleasurable or financially rewarding.

15 Flanagan has defined five domains of QOL, capturing 15 dimensions of life quality. The five domains, and their component dimensions, are physical and material well being (Material well-being and financial security; Health and personal safety), Relations with other people (relations with spouse; Having and rearing children; Relations with parents, siblings, or other

relatives; Relations with friends) Social, community, civic activities (Helping and encouraging others; Participating in local and governmental affairs), Personal development, fulfillment (Intellectual development; Understanding and planning; Occupational role career; Creativity and personal expression), and recreation (Socializing with others; Passive and observational recreational activities; Participating in active recreation). See Flanagan JC,. "A research approach to improving our quality of life." Am Psychol 33:138-147 (1978).

"Health-related quality of life" (HRQL or HRQOL) is an individual's satisfaction or happiness with domains of life insofar as they affect or are affected by "health".

50

In a preferred embodiment, a pharmaceutical agent of the present invention is able to achieve a statistically significant improvement in the expected quality of life, measured according to a commonly accepted measure of QOL, in a treatment group over a control group.

While there is general acceptance of the notion that QOL is important, quantifying QOL is not especially straightforward. Also, QOL can only be measured in humans. Measurements of QOL can be objective (e.g., employment status, marital status, home ownership) or subjective (the subject's opinion of his or her life), or some combination of the two.

A simple approach to measuring subjective QOL is to simply have the subjects rate their overall quality of life on a scale, e.g., of 7 points. One can also use more elaborate measure, such as the Older Adult Health and Mood Questionaire (a 22 item test for assessing depression). Objective QOL can be measured by, e.g., an activities checklist.

There is a relationship between QOL assessment and so-called ADL or IADL measures, which assess the need for assistance.

The Katz Index of Independence in Activities of Daily Living (Katz ADL) measures adequacy of independent performance of bathing, dressing, toileting, transferring, continence, and feeding. See Katz, S., "Assessing Self-Maintenance: Activities of Daily Living, Mobility and Instrumental Activities of Daily Living, Journal of the American Geriatrics Society, 31(12); 721-726 (1983); Katz S., Down, T.D., Cash, H.R. et al. Progress in the Development of the Index of ADL. Gerontologist, 10: 20-30 (1970).

Performance of a more sophisticated nature is measured by

51

the "Instrumental Activities of Daily Living" (IADL) scale. This inquires into ability to independently use the telephone, shop, prepare food, carry out housekeeping, do laundry, travel locally, take medication and handle finances. See Lawton, MP and Brody, EM, Gerontologist, 9:179-86 (1969).

The 36 question Medical Outcomes Study Short Form (SF-36) (Medical Outcomes Trust, Inc., 20 Park Plaza, Suite 1014, Boston, Massachusetts 02116) assesses eight health concepts: 1) limitations in physical activities because of health problems; 2) limitations in social activities because of physical or emotional problems; 3) limitations in usual role activities because of physical health problems; 4) bodily pain; 5) general mental health (psychological distress and well-being); 6) limitations in usual role activities because of emotional problems; 7) vitality (energy and fatigue); and 8) general health perceptions.

A low score on an ADL, IADL or SF-36 test is likely to be associated with a low QOL, but a high score does not guarantee a high QOL because these tests do not explore performance of "valued activities", only of more basic activities. Nonetheless, these tests can be considered commonly accepted measures of QOL for the purpose of this invention.

Age-Related Diseases

5

10

15

10

5

Age-related (senescent) diseases include certain cancers, atherosclerosis, diabetes (type 2), osteoporosis, hypertension, depression, Alzheimer's, Parkinson's, glaucoma, certain immune system defects, kidney failure, and liver steatosis. In general, they are diseases for which the relative risk (comparing a subpopulation over age 55 to a suitably matched population under age 55) is at least 1.1.

Preferably, the agents of the present invention protect

52

against one or more age-related diseases for at least a subpopulation of mature (post-puberty) adult subjects.

Diabetes

5

10

15

3.0

25

:5

Type II diabetes is of particular interest. A deficiency of insulin in the body results in diabetes mellitus, which affects about 18 million individuals in the United States. It is characterized by a high blood glucose (sugar) level and glucose spilling into the urine due to a deficiency of insulin. As more glucose concentrates in the urine, more water is excreted, resulting in extreme thirst, rapid weight loss, drowsiness, fatigue, and possibly dehydration. Because the cells of the diabetic cannot use glucose for fuel, the body uses stored protein and fat for energy, which leads to a buildup of acid (acidosis) in the blood. If this condition is prolonged, the person can fall into a diabetic coma, characterized by deep labored breathing and fruity-odored breath.

There are two types of diabetes mellitus, Type I and Type II. Type II diabetes is the predominant form found in the Western world; fewer than 8% of diabetic Americans have the type I disease.

Type I diabetes. In Type I diabetes, formerly called juvenile-onset or insulin-dependent diabetes mellitus, the pancreas cannot produce insulin. People with Type I diabetes must have daily insulin injections. But they need to avoid taking too much insulin because that can lead to insulin shock, which begins with a mild hunger. This is quickly followed by sweating, shallow breathing, dizziness, palpitations, trembling, and mental confusion. As the blood sugar falls, the body tries to compensate by breaking down fat and protein to make more sugar. Eventually, low blood sugar leads to a decrease in the sugar supply to the brain, resulting in a loss of consciousness. Eating a sugary food can prevent insulin shock until appropriate medical measures

53

can be taken.

5

10

15

30

:5

0

5

Type I diabetics are often characterized by their low or absent levels of circulating endogenous insulin, i.e., hypoinsulinemia (1). Islet cell antibodies causing damage to the pancreas are frequently present at diagnosis. Injection of exogenous insulin is required to prevent ketosis and sustain life.

Type II diabetes. Type II diabetes, formerly called adult-onset or non-insulin-dependent diabetes mellitus (NIDDM), can occur at any age. The pancreas can produce insulin, but the cells do not respond to it.

Type II diabetes is a metabolic disorder that affects approximately 17 million Americans. It is estimated that another 10 million individuals are "prone" to becoming diabetic. These vulnerable individuals can become resistant to insulin, a pancreatic hormone that signals glucose (blood sugar) uptake by fat and muscle. In order to maintain normal glucose levels, the islet cells of the pancreas produce more insulin, resulting in a condition called hyperinsulinemia. When the pancreas can no longer produce enough insulin to compensate for the insulin resistance, and thereby maintain normal glucose levels, hyperglycemia (elevated blood glucose) results, and type II diabetes is diagnosed.

Early Type II diabetics are often characterized by hyperinsulinemia and resistance to insulin. Late Type II diabetics may be normoinsulinemic or hypoinsulinemic. Type II diabetics are usually not insulin dependent or prone to ketosis under normal circumstances.

Little is known about the disease progression from the normoinsulinemic state to the hyperinsulinemic state, and from the hyperinsulinemic state to the Type II diabetic state.

As stated above, type II diabetes is a metabolic disorder that is characterized by insulin resistance and

54

impaired glucose-stimulated insulin secretion (2,3,4). However, Type II diabetes and atherosclerotic disease are viewed as consequences of having the insulin resistance syndrome (IRS) for many years (5). The current theory of the pathogenesis of Type II diabetes is often referred to as the "insulin resistance/islet cell exhaustion" theory. According to this theory, a condition causing insulin resistance compels the pancreatic islet cells to hypersecrete insulin in order to maintain glucose homeostasis. However, after many years of hypersecretion, the islet cells eventually fail and the symptoms of clinical diabetes are manifested. Therefore, this theory implies that, at some point, peripheral hyperinsulinemia will be an antecedent of Type II diabetes. Peripheral hyperinsulinemia can be viewed as the difference between what is produced by the beta cell minus that which is taken up by the liver. Therefore, peripheral hyperinsulinemia can be caused by increased β cell production, decreased hepatic uptake or some combination of both. It is also important to note that it is not possible to determine the origin of insulin resistance once it is established since the onset of peripheral hyperinsulinemia leads to a condition of global insulin resistance.

Multiple environmental and genetic factors are involved in the development of insulin resistance, hyperinsulinemia and type II diabetes. An important risk factor for the development of insulin resistance, hyperinsulinemia and type II diabetes is obesity, particularly visceral obesity (6,7,8). Type II diabetes exists world-wide, but in developed societies, the prevalence has risen as the average age of the population increases and the average individual becomes more obese.

Role of the Liver in the Development of Diabetes

Insulin stimulates the liver to store glucose in the

55

form of glycogen. A large fraction of glucose absorbed from the small intestine is immediately taken up by hepatocytes, which convert it into the storage polymer glycogen. Hepatic uptake of insulin is a function of the number and efficiency of the liver's insulin receptors, and the factors which affect them are not well understood.

In the liver, insulin activates the enzyme hexokinase, which phosphorylates glucose, trapping it within the cell. Insulin also activates several of the enzymes that are directly involved in glycogen synthesis, including phosphofructokinase and glycogen synthase. However, insulin also acts to inhibit the activity of glucose-6-phosphatase.

When the liver is saturated with glycogen, any additional glucose taken up by hepatocytes is shunted into pathways leading to synthesis of fatty acids, which are exported from the liver as lipoproteins. The lipoproteins are ripped apart in the circulation, providing free fatty acids for use in other tissues, including adipocytes, which use them to synthesize triglyceride.

In the absence of insulin, glycogen synthesis in the liver ceases and enzymes responsible for breakdown of glycogen become active.

As noted above, peripheral hyperinsulinemia can be viewed as the difference between what insulin is produced by the β cell minus that which is taken up by the liver. Therefore, peripheral hyperinsulinemia can be caused by increased β cell production, decreased hepatic uptake or some combination of both.

Effect of Diabetes on the Liver

5

5

Diabetes is associated with nonalcoholic steatohepatitis (NASH), also known as nonalcoholic fatty liver disease (NAFLD). In NASH, fat builds up in the liver and eventually causes scar tissue (cirrhosis of the liver).

Non-alcoholic fatty liver disease (NAFLD) is now recognized as one of the most common causes of liver disease

56

and is estimated to affect 10 to 24% of the general population. The higher prevalence of NAFLD in persons with obesity, hyperinsulinemia or type-II diabetes suggests that diet and insulin resistance may play a pivotal role in the development of this syndrome. NAFLD is a clinicopathologic syndrome with a wide spectrum of liver damage ranging from simple steatosis to steatohepatitis (NASH) to advanced fibrosis and cirrhosis. Hepatic steatosis is caused by lipid accumulation within hepatocytes and is a relatively benign condition. However steatosis combined with necroinflammatory activity may progress to end-stage liver disease. It appears that the disease progression requires cellular injury and inflammation in a steatotic environment. While the cause of the injury is not understood, it is clear that hepatic apoptosis is a prominent feature of nonalcoholic steatosis as well as other liver diseases. generally Alba, L.M., Lindor, K. (2003) Review article: Non-alcoholic fatty liver disease., Aliment Pharmacol. Ther. 17:977-986; Ludwig, J., Viggiano, T.R., McGill, D.B., Oh, (1980) Nonalcoholic steatohepatitis: Mayo Clinic experiences with a hitherto unnamed disease. Mayo Clin. Proc. 55:434-438; Chitturi, S., Abeygunasekera, S., Farrel, G.C., Holmes-Walker, J., Hui, J.M., Fung, C., Karim, R., Lin, R., Samarasinghe, D., Liddle, C., Weltman, M., George, J. (2002) NASH and insulin resistance: Insulin hypersecretion and specific association with the insulin resistance syndrome. Hepatology 35:373-379; Feldstein, A.E., Canbay, A., Angulo, P., Taniai, M., Burgart, L.J., Lindor, K.D., Gores, G.J. (2003) Hepatocyte apoptosis and fas expression are prominent features of human nonalcoholic steatohepatitis. Gastroenterology 125:437-443; Higuch, H., Gores, G.J. (2003) Mechanisms of liver injury: an overview. Curr. Mol. Med. 3:483-490.

10

15

3.0

:5

Drugs used for the treatment of diabetes, such as Rezulin (troglitazone), can cause liver damage.

57

Diseases Characterized by Accelerated Aging

Several human diseases display some features of accelerated aging. These include Werner's syndrome (classic early-onset progeria), Hutchinson-Gilford syndrome (adult progeria), and Down's syndrome (trisomy 21). Troen, Biology of Aging, Mt. Sinai J. Med., 70(1): 3 (Jan. 2003). Thus, the present invention may be useful in the treatment (curative or ameliorative) of individuals with these diseases.

ΙQ

.5

:0

5

5

Direct and Indirect Utility of Identified Nucleic Acid Sequences and Related Molecules

The mouse or human genes may be used directly. For diagnostic or screening purposes, they (or specific binding fragments thereof) may be labeled and used as hybridization probes. For therapeutic purposes, they (or specific binding fragments thereof) may be used as antisense reagents to inhibit the expression of the corresponding gene, or of a sufficiently homologous gene of another species.

If the database DNA appears to be a full-length cDNA or gDNA, that is, that it encodes an entire, functional, naturally occurring protein, then it may be used in the expression of that protein. Likewise, if the corresponding human gene is known in full-length, it may be used to express the human protein. Such expression may be in cell culture, with the protein subsequently isolated and administered exogenously to subjects who would benefit therefrom, or in vivo, i.e., administration by gene therapy. Naturally, any DNA encoding the same protein may be used for the same purpose, or a DNA which encodes a fragment or a mutant of that naturally occurring protein which retains the desired activity may be used for the purpose of producing the active fragment or mutant. The encoded protein of course has utility therapeutically and, in labeled or immobilized form, diagnostically.

The genes may also be used indirectly, that is, to

5.8

identify other useful DNAs, proteins, or other molecules. We have attempted to determine whether the mouse genes disclosed herein have significant similarity to any known human DNA, and whether, in any of the six possible combinations of reference frame and strand, they encode a protein similar to a known human protein. If so, then it follows that the known human protein, and DNAs encoding that protein, may be used in a similar manner. In addition, if the known human protein is known to have additional homologues, then those homologous proteins, and DNAs encoding them, may be used in a similar manner.

There thus are several ways that a human protein homologue of interest can be identified by database searching, including:

5

LO

5

:0

- 1) a DNA->DNA (BlastN) search for human database DNAs closely related to the mouse gene identifies a known human gene, and the sequence of the human protein is deduced by the Genetic Code;
- 2) a DNA->Protein (BlastX) search for human database proteins closely related to the translated DNA of the mouse gene identifies a known human protein; and
- 3) the sequence of the mouse protein is known or is deduced by the Genetic Code, and a Protein->Protein (BlastP) search for closely related database proteins identifies a known human protein.

Once a known human gene is identified, it may be used in further BlastN or BlastX searches to identify other human genes or proteins. Once a known human protein is identified, it may be used in further BlastP searches to identify other human proteins. Searches may also take cognizance, intermediately, of known genes and proteins

59

other than mouse or human ones, e.g., use the mouse sequence to identify a known rat sequence and then the rat sequence to identify a human one.

If we have identified a mouse gene (gDNA or cDNA), and it encodes a mouse protein which appears similar to a human protein, then that human protein may be used (especially in humans) for purposes analogous to the proposed use of the mouse protein in mice. Moreover, a specific binding fragment of an appropriate strand of the corresponding human gene (gDNA or cDNA) could be labeled and used as a hybridization probe (especially against samples of human mRNA or cDNA).

5

10

15

20

25

30

35

In determining whether the disclosed genes (gDNA or cDNA) have significant similarities to known DNAs (and their translated AA sequences to known proteins), one would generally use the disclosed gene as a query sequence in a search of a sequence database. The results of several such searches are set forth in the Examples. Such results are dependent, to some degree, on the search parameters. Preferred parameters are set forth in Example 1. The results are also dependent on the content of the database. While the raw similarity score of a particular target (database) sequence will not vary with content (as long as it remains in the database), its informational value (in bits), expected value, and relative ranking can change. Generally speaking, the changes are small.

It will be appreciated that the nucleic acid and protein databases keep growing. Hence a later search may identify high scoring target sequences which were not uncovered by an earlier search because the target sequences were not previously part of a database.

Hence, in a preferred embodiment, the cognate DNAs and proteins include not only those set forth in the examples,

60

but those which would have been highly ranked (top ten, more preferably top three, even more preferably top two, most preferably the top one) in a search run with the same parameters on the date of filing of this application.

5

If the mouse or human database DNA appears to be a partial DNA (that is, partial relative to a cDNA or gDNA encoding the whole naturally occurring protein), it may be used as a hybridization probe to isolate the full-length DNA. If the partial DNA encodes a biologically functional fragment of the cognate protein, it may be used in a manner similar to the full length DNA, i.e., to produce the functional fragment.

15

10

If we have indicated that an antagonist of a protein or other molecule is useful, then such an antagonist may be obtained by preparing a combinatorial library, as described below, of potential antagonists, and screening the library members for binding to the protein or other molecule in question. The binding members may then be further screened for the ability to antagonize the biological activity of the target. The antagonists may be used therapeutically, or, in suitably labeled or immobilized form, diagnostically.

25

20

If the mouse or human database DNA is related to a known protein, then substances known to interact with that protein (e.g., agonists, antagonists, substrates, receptors, second messengers, regulators, and so forth), and binding molecules which bind them, are also of utility. Such binding molecules can likewise be identified by screening a combinatorial library.

30

35

Isolation of Full Length DNAs Using Partial DNAs as probes

If it is determined that a DNA of the present invention is a partial DNA, and the cognate full length DNA is not listed in a sequence database, the available DNA may be used as a hybridization probe to isolate the full-length DNA from

61

a suitable DNA library (cDNA or gDNA).

Stringent hybridization conditions are appropriate, that is, conditions in which the hybridization temperature is 5-10 deg. C. below the Tm of the DNA as a perfect duplex.

5

10

15

20

25

30

35

Identification and Isolation of Homologous Genes Using a DNA Probe

It may be that the sequence databases available do not include the sequence of any homologous gene (gDNA or cDNA), or at least of the homologous gene for a species of interest. However, given the DNAs set forth above, one may readily obtain the homologous gene.

The possession of one DNA (the "starting DNA") greatly facilitates the isolation of homologous DNAs. If only a partial DNA is known, this partial DNA may first be used as a probe to isolate the corresponding full length DNA for the same species, and that the latter may be used as the starting DNA in the search for homologous DNAs.

The starting DNA, or a fragment thereof, is used as a hybridization probe to screen a cDNA or genomic DNA library for clones containing inserts which encode either the entire homologous protein, or a recognizable fragment thereof. The minimum length of the hybridization probe is dictated by the need for specificity. If the size of the library in bases is L, and the GC content is 50%, then the probe should have a length of at least l, where $L=4^{1}$. This will yield, on average, a single perfect match in random DNA of L bases. The human cDNA library is about 10^{8} bases and the human genomic DNA library is about 10^{10} bases.

The library is preferably derived from an organism which is known, on biochemical evidence, to produce a homologous protein, and more preferably from the genomic DNA or mRNA of cells of that organism which are likely to be relatively high producers of that protein. A cDNA library (which is derived from an mRNA library) is especially preferred.

62

If the organism in question is known to have substantially different codon preferences from that of the organism whose relevant cDNA or genomic DNA is known, a synthetic hybridization probe may be used which encodes the same amino acid sequence but whose codon utilization is more similar to that of the DNA of the target organism. Alternatively, the synthetic probe may employ inosine as a substitute for those bases which are most likely to be divergent, or the probe may be a mixed probe which mixes the codons for the source DNA with the preferred codons (encoding the same amino acid) for the target organism.

10

15

3,0

25

; Ó

By routine methods, the Tm of a perfect duplex of starting DNA is determined. One may then select a hybridization temperature which is sufficiently lower than the perfect duplex Tm to allow hybridization of the starting DNA (or other probe) to a target DNA which is divergent from the starting DNA. A 1% sequence divergence typically lowers the Tm of a duplex by 1-2°C, and the DNAs encoding homologous proteins of different species typically have sequence identities of around 50-80%. Preferably, the library is screened under conditions where the temperature is at least 20°C., more preferably at least 50°C., below the perfect duplex Tm. Since salt reduces the Tm, one ordinarily would carry out the search for DNAs encoding highly homologous proteins under relatively <u>low</u> salt hybridization conditions, e.g., <1M NaCl. The higher the salt concentration, and/or the lower the temperature, the greater the sequence divergence which is tolerated.

For the use of probes to identify homologous genes in other species, see, e.g., Schwinn, et al., J. Biol. Chem., 265:8183-89 (1990) (hamster 67-bp cDNA probe vs. human leukocyte genomic library; human 0.32kb DNA probe vs. bovine brain cDNA library, both with hybridization at 42°C in 6xSSC); Jenkins et al., J. Biol. Chem., 265:19624-31 (1990) (Chicken 770-bp cDNA probe vs. human genomic libraries; hybridization at 40°C in 50% formamide and 5xSSC); Murata et

63

al., J. Exp. Med., 175:341-51 (1992) (1.2-kb mouse cDNA probe v. human eosinophil cDNA library; hybridization at 65°C in 6xSSC); Guyer et al., J. Biol. Chem., 265:17307-17 (1990) (2.95-kb human genomic DNA probe vs. porcine genomic DNA library; hybridization at 42°C in 5xSSC). The conditions set forth in these articles may each be considered suitable for the purpose of isolating homologous genes.

Corresponding (Homologous) Proteins and DNAs

In the case of a gene chip, the manufacturer of the gene chip determines which DNA to place at each position on the chip. This DNA may correspond in sequence to a genomic DNA, a cDNA, or a fragment of genomic or cDNA, and may be natural, synthetic or partially natural and partially synthetic in origin. The manufacturer of the gene chip will normally identify the DNA for a mouse gene chip as corresponding to a particular mouse gene, in which case it will be assumed that the alignments of chip DNA to mouse gene satisfies the correspondence (homology) criteria of the invention.

Usually, the gene chip manufacturer will provide a sequence database accession number for the mouse DNA. If so, to identify the corresponding mouse protein, we will first inspect the database record for that mouse DNA. Often, the mouse protein accession number will appear in that record or in a linked record. If it doesn't, the corresponding mouse protein can be identified by performing a BlastX search on a mouse protein database with the mouse database DNA sequence as the query sequence. Even if the protein sequence is not in the database, if the DNA sequence comprises a full-length coding sequence, the corresponding protein can be identified by translating the coding sequence in accordance with the Genetic Code.

5

5

LO

رL5

30

:5

0

64

corresponding (homologous) to a gene chip DNA if it is identified as corresponding (homologous) to the mouse gene (gDNA or cDNA, whole or partial) identified by the gene chip manufacturer as corresponding (homologous) to that gene chip DNA.

In turn, it is identifiable as corresponding (homologous) to said identified mouse gene, if

- (1) it can be aligned by BlastX directly to that mouse gene, and/or
- (2) it is encoded by a human gene, or can be aligned to a human gene by BlastX, which in turn can be aligned by BlastN to said mouse gene and/or
- (3) it can be aligned by BlastP to a mouse protein, the latter being encoded by said mouse gene, or aligned to said mouse gene BlastX,

where any alignment by BlastN, BlastP or BlastX is in accordance with the default parameters set forth below, and the expected value (E) of each alignment (the probability that such an alignment would have occurred by chance alone) is less than e-10. (Note that because this is a negative exponent, a value such as e-50 is less than e-10.)

A human gene is corresponding (homologous) to a mouse gene chip DNA, and hence to said identified mouse gene (or cDNA) and protein, if it encodes a corresponding (homologous) human protein as defined above, or it can be aligned by BlastN to said mouse gene.

Desirably, two or all three of these conditions (1)-(3) are satisfied for the corresponding (homologous) human genes and proteins.

65

Preferably, for at least one of conditions (1)-(3), the E value is less than e-50, more preferably less than e-60, still more preferably less than e-70, even more preferably less than e-80, considerably more preferably less than e-90, and most preferably less than e-100. Desirably, it is true for two or even all three of these conditions.

5

LO

.5

:0

.5

0

5

In constructing Master table 1, we generally used a BlastX (mouse gene vs. human protein) alignment E value cutoff of e-50. However, if there were no human proteins with that good an alignment to the mouse DNA in question, or if there were other reasons for including a particular human protein (e.g., a known functionality supportive of the observed differential cognate mouse protein expression), then a human protein with a score worse (i.e., higher) than e-50 may appear in Master Table 1.

BlastN and BlastX report very low expected values as "0.0". This does not truly mean that the expected value is exactly zero (since any alignment could occur by chance), but merely that it is so infinitesimal that it is not reported. The documentation does not state the cutoff value, alignments with explicit E values as low as e-178 (624 bits) have been reported as such, while a score of 636 bits was reported as "0.0".

If the manufacturer of the gene chip identifies the gene chip DNA as corresponding to an EST, or other DNA which is not a full-length mouse gene or cDNA, a longer (possibly full length) mouse gene or cDNA may be identified by a BlastN search of the mouse DNA database. Alternatively, the identified DNA may be used to conduct a BlastN search of a human DNA database, or a BlastX search of a mouse or human protein database.

Thus, more generally, a human protein can be said to be

66

identifiable as corresponding (homologous) to a gene chip DNA, or to a DNA identified by the manufacturer as corresponding to that gene chip DNA, if

- (1') it can be aligned directly to the gene chip or corresponding manufacturer identified DNA by BlastX. and/or
- (2') it can be aligned to a human gene/cDNA by BlastX, whose genomic DNA (gDNA) or cDNA (DNA complementary to messenger RNA) in turn can be aligned to the gene chip or corresponding manufacturer identified DNA by BlastN, and/or

ĹΟ

L5

310

25

ijΰ

.5

- (3') it can be aligned to a mouse gene/cDNA by BlastX, whose gDNA or cDNA in turn can be aligned to the gene chip or corresponding manufacturer identified DNA by BlastN, and/or
- (4') it can be aligned to a mouse protein by BlastP, which in turn can be aligned to the gene chip or corresponding manufacturer identified DNA by BlastX, and/or
- (5') it can be aligned to a mouse protein by BlastP, which in turn can be aligned to a mouse gene/cDNA by BlastX, whose gDNA or cDNA can in turn be aligned to the gene chip or corresponding manufacturer identified DNA by BlastN;

where any alignment by BlastN, BlastP, or BlastX is in accordance with the default parameters set forth below, and the expected value (E) of each alignment (the probability that such an alignment would have occurred by chance alone) is less than e-10. (Note that because this is a negative exponent, a value such as e-50 is less than e-10.)

Preferably, two, three, four or all five of conditions (1')-(5') are satisfied.

Preferably, for at least one of conditions (1')-(5'), for at least the final alignment (i.e., vs. the human

67

protein), the E value is less than e-50, more preferably less than e-60, , still more preferably less than e-70, even more preferably less than e-80, considerably more preferably less than e-90, and most preferably less than e-100.

5

10

15

30

!5

0

Desirably, one or more of these standards of preference are met for two, three, four or all five of conditions (1')-(5'). In particular, for those conditions in which the gene chip or corresponding manufacturer identified DNA is indirectly connected to the human protein by virtue of two or more successive alignments, the E value is preferably, so limited for all of said alignments in the connecting chain.

A human gene corresponds (is homologous) to a gene chip DNA or manufacturer identified corresponding DNA if it encodes a corresponding (homologous) human protein as defined above, or if it can be aligned either directly to that DNA, or indirectly through a mouse gene which can be aligned to said DNA, according to the conditions set forth above.

Master table 1 assembles a list of human protein corresponding (homologous) to each of the mouse DNAs/proteins identified as related to the chip DNA. These human proteins form a set and can be given a percentile rank, with respect to E value, within that set. The human proteins of the present invention preferably are those scorers with a percentile rank of at least 50%, more preferably at least 60%, still more preferably at least 70%, even more preferably at least 80%, and most preferably at least 90%.

For each mouse gene in Master Table 1, there is a particular human protein which provides the best alignment match as measured by BlastX, i.e., the human protein with the best score (lowest e-value). These human proteins form a subset of the set above and can be given a percentile rank within that subset, e.g., the human proteins with scores in

68

the top 10% of that subset have a percentile rank of 90% or higher.

The human proteins of the present invention preferably are those best scorer subset proteins with a percentile rank within the subset of at least 50%, more preferably at least 60%, still more preferably at least 70%, even more preferably at least 80%, and most preferably at least 90%.

BlastN and BlastX report very low expected values as "0.0". This does not truly mean that the expected value is exactly zero (since any alignment could occur by chance), but merely that it is so infinitesimal that it is not reported. The documentation does not state the cutoff value, but alignments with explicit E values as low as e-178 (624 bits) have been reported as nonzero values, while a score of 636 bits was reported as "0.0".

Functionally homologous human proteins are also of interest. A human protein may be said to be functionally homologous to the mouse gene if the human protein has at least one biological activity in common with the mouse protein encoded by said mouse gene.

The human proteins of interest also include those that are substantially and/or conservatively identical (as defined below) to the homologous and/or functionally homologous human proteins defined above.

Degree of Differential Expression

5

10

15

20

25

30

35

The degree of differential expression may be expressed as the ratio of the higher expression level to the lower expression level. Preferably, this is at least 2-fold, and more preferably, it is higher, such as at least 3-fold, at least 4-fold, at least 5-fold, at least 6-fold, at least 7-fold, at least 8-fold, at least 9-fold, Or at least 10-fold.

Most preferably, the human protein of interest

69

corresponds to a mouse gene for which the degree of differential expression places it among the top 10% of the mouse genes in the appropriate subtable.

Relevance of Favorable and Unfavorable Genes

5

.0

.5

20

:5

10

;5

If a gene is down-regulated in more favored mammals, or up-regulated in less favored mammals, (i.e., an "unfavorable gene") then several utilities are apparent.

First, the complementary strand of the gene, or a portion thereof, may be used in labeled form as a hybridization probe to detect messenger RNA and thereby monitor the level of expression of the gene in a subject. Elevated levels are indicative of progression, or propensity to progression, to a less favored state, and clinicians may take appropriate preventative, curative or ameliorative action.

Secondly, the messenger RNA product (or equivalent cDNA), the protein product, or a binding molecule specific for that product (e.g., an antibody which binds the product), or a downstream product which mediates the activity (e.g., a signaling intermediate) or a binding molecule (e.g., an antibody) therefor, may be used, preferably in labeled or immobilized form, as an assay reagent in an assay for said nucleic acid product, protein product, or downstream product (e.g., a signaling intermediate). Again, elevated levels are indicative of a present or future problem.

Thirdly, an agent which down-regulates expression of the gene may be used to reduce levels of the corresponding protein and thereby inhibit further damage. This agent could inhibit transcription of the gene in the subject, or translation of the corresponding messenger RNA. Possible inhibitors of transcription and translation include antisense molecules and repressor molecules. The agent could also inhibit a post-translational modification (e.g., glycosylation, phosphorylation, cleavage, GPI attachment)

70

required for activity, or post-translationally modify the protein so as to inactivate it. Or it could be an agent which down- or up-regulated a positive or negative regulatory gene, respectively.

5

LΟ

:10

:5

Fourthly, an agent which is an antagonist of the messenger RNA product or protein product of the gene, or of a downstream product through which its activity is manifested (e.g., a signaling intermediate), may be used to inhibit its activity.

This antagonist could be an antibody, a peptide, a peptide, a nucleic acid, a peptide nucleic acid (PNA) oligomer, a small organic molecule of a kind for which a combinatorial library exists (e.g., a benzodiazepine), etc. An antagonist is simply a binding molecule which, by binding, reduces or abolishes the undesired activity of its target. The antagonist, if not an oligomeric molecule, is preferably less than 1000 daltons, more preferably less than 500 daltons.

Fifthly, an agent which degrades, or abets the degradation of, that messenger RNA, its protein product or a downstream product which mediates its activity (e.g., a signaling intermediate), may be used to curb the effective period of activity of the protein.

If a gene is <u>up</u>-regulated in more favored mammals, or <u>down</u>-regulated in less favored animals then the utilities are converse to those stated above.

First, the complementary strand of the gene, or a portion thereof, may be used in labeled form as a hybridization probe to detect messenger RNA and thereby monitor the level of expression of the gene in a subject. Depressed levels are indicative of damage, or possibly of a propensity to damage, and clinicians may take appropriate preventative, curative or ameliorative action.

Secondly, the messenger RNA product, the equivalent cDNA, protein product, or a binding molecule specific for those products, or a downstream product, or a signaling

71

intermediate, or a binding molecule therefor, may be used, preferably in labeled or immobilized form, as an assay reagent in an assay for said protein product or downstream product. Again, depressed levels are indicative of a present or future problem.

Thirdly, an agent which up-regulates expression of the gene may be used to increase levels of the corresponding protein and thereby inhibit further progression to a less favored state. By way of example, it could be a vector which carries a copy of the gene, but which expresses the gene at higher levels than does the endogenous expression system. Or it could be an agent which up- or down-regulates a positive or negative regulatory gene.

Fourthly, an agent which is an agonist of the protein product of the gene, or of a downstream product through which its activity (of inhibition of progression to a less favored state) is manifested, or of a signaling intermediate may be used to foster its activity.

Fifthly, an agent which inhibits the degradation of that protein product or of a downstream product or of a signaling intermediate may be used to increase the effective period of activity of the protein.

Mutant Proteins

The present invention also contemplates mutant proteins (peptides) which are substantially identical (as defined below) to the parental protein (peptide). In general, the fewer the mutations, the more likely the mutant protein is to retain the activity of the parental protein. The effect of mutations is usually (but not always) additive. Certain individual mutations are more likely to be tolerated than others.

A protein is more likely to tolerate a mutation which

(a) is a substitution rather than an insertion or deletion:

- (b) is an insertion or deletion at the terminus, rather than internally, or, if internal, is at a domain boundary, or a loop or turn, rather than in an alpha helix or beta strand;
- (c) affects a surface residue rather than an interior residue:
- (d) affects a part of the molecule distal to the binding site;
- (e) is a substitution of one amino acid for another of similar size, charge, and/or hydrophobicity, and does not destroy a disulfide bond or other crosslink; and
- (f) is at a site which is subject to substantial variation among a family of homologous proteins to which the protein of interest belongs.

These considerations can be used to design functional mutants.

Surface vs. Interior Residues

5

10

15

30

35

30

Charged amino acid residues almost always lie on the surface of the protein. For uncharged residues, there is less certainty, but in general, hydrophilic residues are partitioned to the surface and hydrophobic residues to the interior. Of course, for a membrane protein, the membrane-spanning segments are likely to be rich in hydrophobic residues.

Surface residues may be identified experimentally by various labeling techniques, or by 3-D structure mapping techniques like X-ray diffraction and NMR. A 3-D model of a homologous protein can be helpful.

Binding Site Residues

Residues forming the binding site may be identified by (1) comparing the effects of labeling the surface residues before and after complexing the protein to its target, (2) labeling the binding site directly with affinity ligands,

PCT/US2004/017322

73

(3) fragmenting the protein and testing the fragments for binding activity, and (4) systematic mutagenesis (e.g., alanine-scanning mutagenesis) to determine which mutants destroy binding. If the binding site of a homologous protein is known, the binding site may be postulated by analogy.

Protein libraries may be constructed and screened that a large family (e.g., 10^8) of related mutants may be evaluated simultaneously.

Hence, the mutations are preferably conservative modifications as defined below.

"Substantially Identical"

A mutant protein (peptide) is substantially identical to a reference protein (peptide) if (a) it has at least 10% of a specific binding activity or a non-nutritional biological activity of the reference protein, and (b) is at least 50% identical in amino acid sequence to the reference protein (peptide). It is "substantially structurally identical" if condition (b) applies, regardless of (a).

Percentage amino acid identity is determined by aligning the mutant and reference sequences according to a rigorous dynamic programming algorithm which globally aligns their sequences to maximize their similarity, the similarity being scored as the sum of scores for each aligned pair according to an unbiased PAM250 matrix, and a penalty for each internal gap of -12 for the first null of the gap and -4 for each additional null of the same gap. The percentage identity is the number of matches expressed as a percentage of the adjusted (i.e., counting inserted nulls) length of the reference sequence.

A mutant DNA sequence is substantially identical to a reference DNA sequence if they are structural sequences, and encoding mutant and reference proteins which are substantially identical as described above.

If instead they are regulatory sequences, they are

74

substantially identical if the mutant sequence has at least 10% of the regulatory activity of the reference sequence, and is at least 50% identical in nucleotide sequence to the reference sequence. Percentage identity is determined as for proteins except that matches are scored +5, mismatches -4, the gap open penalty is -12, and the gap extension penalty (per additional null) is -4.

More preferably, the sequence is not merely substantially identical, but rather is at least 51%, 66%, 75%, 80%, 85%, 90%, 95%, 96%, 97%, 98% or 99% identical in sequence to the reference sequence.

DNA sequences may also be considered "substantially identical" if they hybridize to each other under stringent conditions, i.e., conditions at which the Tm of the heteroduplex of the one strand of the mutant DNA and the more complementary strand of the reference DNA is not in excess of 10°C. less than the Tm of the reference DNA homoduplex. Typically this will correspond to a percentage identity of 85-90%.

"Conservative Modifications"

LO

15

10

"Conservative modifications" are defined as

- (a) conservative substitutions of amino acids as hereafter defined; or
- (b) single or multiple insertions (extension) or deletions (truncation) of amino acids at the termini.

Conservative modifications are preferred to other modifications. Conservative substitutions are preferred to other conservative modifications.

"Semi-Conservative Modifications" are modifications which are not conservative, but which are (a) semi-conservative substitutions as hereafter defined; or (b) single or multiple insertions or deletions internally, but at interdomain boundaries, in loops or in other segments of relatively high mobility. Semi-conservative modifications

75

are preferred to nonconservative modifications. Semiconservative substitutions are preferred to other semiconservative modifications.

5

5

0

5

5

Non-conservative substitutions are preferred to other non-conservative modifications.

The term "conservative" is used here in an <u>a priori</u> sense, i.e., modifications which would be <u>expected</u> to preserve 3D structure and activity, based on analysis of the naturally occurring families of homologous proteins and of past experience with the effects of deliberate mutagenesis, rather than <u>post facto</u>, a modification already known to conserve activity. Of course, a modification which is conservative <u>a priori</u> may, and usually is, also conservative <u>post facto</u>.

Preferably, except at the termini, no more than about five amino acids are inserted or deleted at a particular locus, and the modifications are outside regions known to contain binding sites important to activity.

Preferably, insertions or deletions are limited to the termini.

A conservative substitution is a substitution of one amino acid for another of the same exchange group, the exchange groups being defined as follows

- I Gly, Pro, Ser, Ala (Cys) (and any nonbiogenic, neutral amino acid with a hydrophobicity not exceeding that of the aforementioned a.a.'s)
- II Arg, Lys, His (and any nonbiogenic, positivelycharged amino acids)
- IV Leu, Ile, Met, Val (Cys) (and any nonbiogenic, aliphatic, neutral amino acid with a hydrophobicity too high for I above)
- V Phe, Trp, Tyr (and any nonbiogenic, aromatic neutral amino acid with a hydrophobicity too high for I above).

76

Note that Cys belongs to both I and IV.

Residues Pro, Gly and Cys have special conformational roles. Cys participates in formation of disulfide bonds. Gly imparts flexibility to the chain. Pro imparts rigidity to the chain and disrupts α helices. These residues may be essential in certain regions of the polypeptide, but substitutable elsewhere.

One, two or three conservative substitutions are more likely to be tolerated than a larger number.

"Semi-conservative substitutions" are defined herein as being substitutions within supergroup I/II/III or within supergroup IV/V, but not within a single one of groups I-V. They also include replacement of any other amino acid with alanine. If a substitution is not conservative, it preferably is semi-conservative.

"Non-conservative substitutions" are substitutions which are not "conservative" or "semi-conservative".

"Highly conservative substitutions" are a subset of conservative substitutions, and are exchanges of amino acids within the groups Phe/Tyr/Trp, Met/Leu/Ile/Val, His/Arg/Lys, Asp/Glu and Ser/Thr/Ala. They are more likely to be tolerated than other conservative substitutions. Again, the smaller the number of substitutions, the more likely they are to be tolerated.

"Conservatively Identical"

0

A protein (peptide) is conservatively identical to a reference protein (peptide) it differs from the latter, if at all, solely by conservative modifications, the protein (peptide) remaining at least seven amino acids long if the reference protein (peptide) was at least seven amino acids long.

A protein is at least semi-conservatively identical to a reference protein (peptide) if it differs from the latter, if at all, solely by semi-conservative or conservative modifications.

77

A protein (peptide) is nearly conservatively identical to a reference protein (peptide) if it differs from the latter, if at all, solely by one or more conservative modifications and/or a single nonconservative substitution.

It is highly conservatively identical if it differs, if at all, solely by highly conservative substitutions. Highly conservatively identical proteins are preferred to those merely conservatively identical. An absolutely identical protein is even more preferred.

ſΟ

.5

0:

:5

The core sequence of a reference protein (peptide) is the largest single fragment which retains at least 10% of a particular specific binding activity, if one is specified, or otherwise of at least one specific binding activity of the referent. If the referent has more than one specific binding activity, it may have more than one core sequence, and these may overlap or not.

If it is taught that a peptide of the present invention may have a particular similarity relationship (e.g., markedly identical) to a reference protein (peptide), preferred peptides are those which comprise a sequence having that relationship to a core sequence of the reference protein (peptide), but with internal insertions or deletions in either sequence excluded. Even more preferred peptides are those whose entire sequence has that relationship, with the same exclusion, to a core sequence of that reference protein (peptide).

٠0

5

Library

The term "library" generally refers to a collection of chemical or biological entities which are related in origin, structure, and/or function, and which can be screened simultaneously for a property of interest.

Libraries may be classified by how they are constructed

78

(natural vs. artificial diversity; combinatorial vs. noncombinatorial), how they are screened (hybridization, expression, display), or by the nature of the screened library members (peptides, nucleic acids, etc.).

5

0

.5

0

5

In a "natural diversity" library, essentially all of the diversity arose without human intervention. This would be true, for example, of messenger RNA extracted from a nonengineered cell.

In a "synthetic diversity" library, essentially all of the diversity arose deliberately as a result of human intervention. This would be true for example of a combinatorial library; note that a small level of natural diversity could still arise as a result of spontaneous mutation. It would also be true of a noncombinatorial library of compounds collected from diverse sources, even if they were all natural products.

In a "non-natural diversity" library, at least some of the diversity arose deliberately through human intervention.

In a "controlled origin" library, the source of the diversity is limited in some way. A limitation might be to cells of a particular individual, to a particular species, or to a particular genus, or, more complexly, to individuals of a particular species who are of a particular age, sex, physical condition, geographical location, occupation and/or familial relationship. Alternatively or additionally, it might be to cells of a particular tissue or organ. Or it could be cells exposed to particular pharmacological, environmental, or pathogenic conditions. Or the library could be of chemicals, or a particular class of chemicals, produced by such cells.

In a "controlled structure" library, the library members are deliberately limited by the production conditions to particular chemical structures. For example, if they are oligomers, they may be limited in length and monomer composition, e.g. hexapeptides composed of the twenty genetically encoded amino acids.

WO 2005/000335

79

Hybridization Library

In a hybridization library, the library members are nucleic acids, and are screened using a nucleic acid hybridization probe. Bound nucleic acids may then be amplified, cloned, and/or sequenced.

Expression Library

5

10

15

3.0

25

0:

. .5

In an expression library, the screened library members are gene expression products, but one may also speak of an underlying library of genes encoding those products. The library is made by subcloning DNA encoding the library members (or portions thereof) into expression vectors (or into cloning vectors which subsequently are used to construct expression vectors), each vector comprising an expressible gene encoding a particular library member, introducing the expression vectors into suitable cells, and expressing the genes so the expression products are produced.

In one embodiment, the expression products are secreted, so the library can be screened using an affinity reagent, such as an antibody or receptor. The bound expression products may be sequenced directly, or their sequences inferred by, e.g., sequencing at least the variable portion of the encoding DNA.

In a second embodiment, the cells are lysed, thereby exposing the expression products, and the latter are screened with the affinity reagent.

In a third embodiment, the cells express the library members in such a manner that they are displayed on the surface of the cells, or on the surface of viral particles produced by the cells. (See display libraries, below).

In a fourth embodiment, the screening is not for the ability of the expression product to bind to an affinity reagent, but rather for its ability to alter the phenotype of the host cell in a particular detectable manner. Here, the screened library members are transformed cells, but

8.0

there is a first underlying library of expression products which mediate the behavior of the cells, and a second underlying library of genes which encode those products.

Display Library

5

<u>!</u>5

:0

:5

In a display library, the library members are each conjugated to, and displayed upon, a support of some kind. The support may be living (a cell or virus), or nonliving (e.g., a bead or plate).

If the support is a cell or virus, display will normally be effectuated by expressing a fusion protein which comprises the library member, a carrier moiety allowing integration of the fusion protein into the surface of the cell or virus, and optionally a lining moiety. In a variation on this theme, the cell coexpresses a first fusion comprising the library member and a linking moiety L1, and a second fusion comprising a linking moiety L2 and the carrier moiety. L1 and L2 interact to associate the first fusion with the second fusion and hence, indirectly, the library member with the surface of the cell or virus.

Soluble Library

In a soluble library, the library members are free in solution. A soluble library may be produced directly, or one may first make a display library and then release the library members from their supports.

Encapsulated Library

In an encapsulated library, the library members are inside cells or liposomes. Generally speaking, encapsulated libraries are used to store the library members for future use; the members are extracted in some way for screening purposes. However, if they differentially affect the phenotype of the cells, they may be screened indirectly by screening the cells.

cDNA Library

5

.0

.5

0

5

0

5

A cDNA library is usually prepared by extracting RNA from cells of particular origin, fractionating the RNA to isolate the messenger RNA (mRNA has a poly(A) tail, so this is usually done by oligo-dT affinity chromatography), synthesizing complementary DNA (cDNA) using reverse transcriptase, DNA polymerase, and other enzymes, subcloning the cDNA into vectors, and introducing the vectors into cells. Often, only mRNAs or cDNAs of particular sizes will be used, to make it more likely that the cDNA encodes a functional polypeptide.

A cDNA library explores the natural diversity of the transcribed DNAs of cells from a particular source. It is not a combinatorial library.

A cDNA library may be used to make a hybridization library, or it may be used as an (or to make) expression library.

Genomic DNA Library

A genomic DNA library is made by extracting DNA from a particular source, fragmenting the DNA, isolating fragments of a particular size range, subcloning the DNA fragments into vectors, and introducing the vectors into cells.

Like a cDNA library, a genomic DNA library is a natural diversity library, and not, a combinatorial library. A genomic DNA library may be used the same way as a cDNA library.

Synthetic DNA library

A synthetic DNA library may be screened directly (as a hybridization library), or used in the creation of an expression or display library of peptides/proteins.

Combinatorial Libraries

The term "combinatorial library" refers to a library in which the individual members are either systematic or random

82

5

0

5

combinations of a limited set of basic elements, the properties of each member being dependent on the choice and location of the elements incorporated into it. Typically, the members of the library are at least capable of being screened simultaneously. Randomization may be complete or partial; some positions may be randomized and others predetermined, and at random positions, the choices may be limited in a predetermined manner. The members of a combinatorial library may be oligomers or polymers of some kind, in which the variation occurs through the choice of monomeric building block at one or more positions of the oligomer or polymer, and possibly in terms of the connecting linkage, or the length of the oligomer or polymer, too. Or the members may be nonoligomeric molecules with a standard core structure, like the 1,4-benzodiazepine structure, with the variation being introduced by the choice of substituents at particular variable sites on the core structure. members may be nonoligomeric molecules assembled like a jigsaw puzzle, but wherein each piece has both one or more variable moieties (contributing to library diversity) and one or more constant moieties (providing the functionalities for coupling the piece in question to other pieces).

Thus, in a typical combinatorial library, chemical building blocks are at least partially randomly combined into a large number (as high as 10¹⁵) of different compounds, which are then simultaneously screened for binding (or other) activity against one or more targets.

In a "simple combinatorial library", all of the members belong to the same class of compounds (e.g., peptides) and can be synthesized simultaneously. A "composite combinatorial library" is a mixture of two or more simple libraries, e.g., DNAs and peptides, or peptides, peptoids, and PNAs, or benzodiazepines and carbamates. The number of component simple libraries in a composite library will, of course, normally be smaller than the average number of members in each simple library, as otherwise the advantage

83

of a library over individual synthesis is small.

5

0

5

0

5

)

Libraries of thousands, even millions, of random oligopeptides have been prepared by chemical synthesis (Houghten et al., Nature, 354:84-6(1991)), or gene expression (Marks et al., J Mol Biol, 222:581-97(1991)), displayed on chromatographic supports (Lam et al., Nature, 354:82-4(1991)), inside bacterial cells (Colas et al., Nature, 380:548-550(1996)), on bacterial pili (Lu, Bio/Technology, 13:366-372(1990)), or phage (Smith, Science, 228:1315-7(1985)), and screened for binding to a variety of targets including antibodies (Valadon et al., J Mol Biol, 261:11-22(1996)), cellular proteins (Schmitz et al., J Mol Biol, 260:664-677(1996)), viral proteins (Hong and Boulanger, Embo J, 14:4714-4727(1995)), bacterial proteins (Jacobsson and Frykberg, Biotechniques, 18:878-885(1995)), nucleic acids (Cheng et al., Gene, 171:1-8(1996)), and plastic (Siani et al., J Chem Inf Comput Sci, 34:588-593 (1994)).

Libraries of proteins (Ladner, USP 4,664,989), peptoids (Simon et al., Proc Natl Acad Sci U S A, 89:9367-71(1992)), nucleic acids (Ellington and Szostak, Nature, 246:818(1990)), carbohydrates, and small organic molecules (Eichler et al., Med Res Rev, 15:481-96(1995)) have also been prepared or suggested for drug screening purposes.

The first combinatorial libraries were composed of peptides or proteins, in which all or selected amino acid positions were randomized. Peptides and proteins can exhibit high and specific binding activity, and can act as catalysts. In consequence, they are of great importance in biological systems.

Nucleic acids have also been used in combinatorial libraries. Their great advantage is the ease with which a nucleic acid with appropriate binding activity can be amplified. As a result, combinatorial libraries composed of nucleic acids can be of low redundancy and hence, of high diversity.

84

There has also been much interest in combinatorial libraries based on small molecules, which are more suited to pharmaceutical use, especially those which, like benzodiazepines, belong to a chemical class which has already yielded useful pharmacological agents. The techniques of combinatorial chemistry have been recognized as the most efficient means for finding small molecules that act on these targets. At present, small molecule combinatorial chemistry involves the synthesis of either pooled or discrete molecules that present varying arrays of functionality on a common scaffold. These compounds are grouped in libraries that are then screened against the target of interest either for binding or for inhibition of biological activity.

)

j

The size of a library is the number of molecules in it. The simple diversity of a library is the number of unique structures in it. There is no formal minimum or maximum diversity. If the library has a very low diversity, the library has little advantage over just synthesizing and screening the members individually. If the library is of very high diversity, it may be inconvenient to handle, at least without automatizing the process. The simple diversity of a library is preferably at least 10, 10E2, 10E3, 10E4, 10E6, 10E7, 10E8 or 10E9, the higher the better under most circumstances. The simple diversity is usually not more than 10E15, and more usually not more than 10E10.

The average sampling level is the size divided by the simple diversity. The expected average sampling level must be high enough to provide a reasonable assurance that, if a given structure were expected, as a consequence of the library design, to be present, that the actual average sampling level will be high enough so that the structure, if satisfying the screening criteria, will yield a positive result when the library is screened. Thus, the preferred average sampling level is a function of the detection limit, which in turn is a function of the strength of the signal to

85

PCT/US2004/017322

be screened.

5 .

.0

.5

0

5

5

WO 2005/000335

There are more complex measures of diversity than simple diversity. These attempt to take into account the degree of structural difference between the various unique sequences. These more complex measures are usually used in the context of small organic compound libraries, see below.

The library members may be presented as solutes in solution, or immobilized on some form of support. In the latter case, the support may be living (cell, virus) or nonliving (bead, plate, etc.). The supports may be separable (cells, virus particles, beads) so that binding and nonbinding members can be separated, or nonseparable (plate). In the latter case, the members will normally be placed on addressable positions on the support. The advantage of a soluble library is that there is no carrier moiety that could interfere with the binding of the members to the support. The advantage of an immobilized library is that it is easier to identify the structure of the members which were positive.

When screening a soluble library, or one with a separable support, the target is usually immobilized. When screening a library on a nonseparable support, the target will usually be labeled.

Oligonucleotide Libraries

An oligonucleotide library is a combinatorial library, at least some of whose members are single-stranded oligonucleotides having three or more nucleotides connected by phosphodiester or analogous bonds. The oligonucleotides may be linear, cyclic or branched, and may include non-nucleic acid moieties. The nucleotides are not limited to the nucleotides normally found in DNA or RNA. For examples of nucleotides modified to increase nuclease resistance and chemical stability of aptamers, see Chart 1 in Osborne and Ellington, Chem. Rev., 97: 349-70 (1997). For screening of RNA, see Ellington and Szostak, Nature, 346: 818-22 (1990).

86

There is no formal minimum or maximum size for these oligonucleotides. However, the number of conformations which an oligonucleotide can assume increases exponentially with its length in bases. Hence, a longer oligonucleotide is more likely to be able to fold to adapt itself to a protein surface. On the other hand, while very long molecules can be synthesized and screened, unless they provide a much superior affinity to that of shorter molecules, they are not likely to be found in the selected population, for the reasons explained by Osborne and Ellington (1997). Hence, the libraries of the present invention are preferably composed of oligonucleotides having a length of 3 to 100 bases, more preferably 15 to 35 bases. The oligonucleotides in a given library may be of the same or of different lengths.

5

0

Oligonucleotide libraries have the advantage that libraries of very high diversity (e.g., 10¹⁵) are feasible, and binding molecules are readily amplified in vitro by polymerase chain reaction (PCR). Moreover, nucleic acid molecules can have very high specificity and affinity to targets.

In a preferred embodiment, this invention prepares and screens oligonucleotide libraries by the SELEX method, as described in King and Famulok, Molec. Biol. Repts., 20: 97-107 (1994); L. Gold, C. Tuerk. Methods of producing nucleic acid ligands, US#5595877; Oliphant et al. Gene 44:177 (1986).

The term "aptamer" is conferred on those oligonucleotides which bind the target protein. Such aptamers may be used to characterize the target protein, both directly (through identification of the aptamer and the points of contact between the aptamer and the protein) and indirectly (by use of the aptamer as a ligand to modify the chemical reactivity of the protein).

In a classic oligonuclotide, each nucleotide (monomeric unit) is composed of a phosphate group, a sugar moiety, and

87

either a purine or a pyrimidine base. In DNA, the sugar is deoxyribose and in RNA it is ribose. The nucleotides are linked by 5'-3' phosphodiester bonds.

The deoxyribose phosphate backbone of DNA can be modified to increase resistance to nuclease and to increase penetration of cell membranes. Derivatives such as mono- or dithiophosphates, methyl phosphonates, boranophosphates, formacetals, carbamates, siloxanes, and dimethylenethio-sulfoxideo-and-sulfono-linked species are known in the art.

Peptide Library

5

10

15

20

25

30

35

A peptide is composed of a plurality of amino acid residues joined together by peptidyl (-NHCO-) bonds. A biogenic peptide is a peptide in which the residues are all genetically encoded amino acid residues; it is not necessary that the biogenic peptide actually be produced by gene expression.

Amino acids are the basic building blocks with which peptides and proteins are constructed. Amino acids possess both an amino group (-NH $_2$) and a carboxylic acid group (-COOH). Many amino acids, but not all, have the alpha amino acid structure NH $_2$ -CHR-COOH, where R is hydrogen, or any of a variety of functional groups.

Twenty amino acids are genetically encoded: Alanine, Arginine, Asparagine, Aspartic Acid, Cysteine, Glutamic Acid, Glutamine, Glycine, Histidine, Isoleucine, Leucine, Lysine, Methionine, Phenylalanine, Proline, Serine, Threonine, Tryptophan, Tyrosine, and Valine. Of these, all save Glycine are optically isomeric, however, only the L-form is found in humans. Nevertheless, the D-forms of these amino acids do have biological significance; D-Phe, for example, is a known analgesic.

Many other amino acids are also known, including: 2-Aminoadipic acid; 3-Aminoadipic acid; beta-Aminopropionic acid; 2-Aminobutyric acid; 4-Aminobutyric acid (Piperidinic

88

acid);6-Aminocaproic acid; 2-Aminoheptanoic acid; 2-Aminoisobutyric acid, 3-Aminoisobutyric acid; 2'-Aminopimelic acid; 2,4-Diaminobutyric acid; Desmosine; 2,2'-Diaminopimelic acid; 2,3-Diaminopropionic acid; N-Ethylglycine; N-Ethylasparagine; Hydroxylysine; allo-Hydroxylysine; 3-Hydroxyproline; 4-Hydroxyproline; Isodesmosine; allo-Isoleucine; N-Methylglycine (Sarcosine); N-Methylisoleucine; N-Methylvaline; Norvaline; Norleucine; and Ornithine.

5

10

15

20

30

35

Peptides are constructed by condensation of amino acids and/or smaller peptides. The amino group of one amino acid (or peptide) reacts with the carboxylic acid group of a second amino acid (or peptide) to form a peptide (-NHCO-) bond, releasing one molecule of water. Therefore, when an amino acid is incorporated into a peptide, it should, technically speaking, be referred to as an amino acid residue. The core of that residue is the moiety which excludes the -NH and -CO linking functionalities which connect it to other residues. This moiety consists of one or more main chain atoms (see below) and the attached side chains.

The main chain moiety of each amino acid consists of the -NH and -CO linking functionalities and a core main chain moiety. Usually the latter is a single carbon atom. However, the core main chain moiety may include additional carbon atoms, and may also include nitrogen, oxygen or sulfur atoms, which together form a single chain. In a preferred embodiment, the core main chain atoms consist solely of carbon atoms.

The side chains are attached to the core main chain atoms. For alpha amino acids, in which the side chain is attached to the alpha carbon, the C-1, C-2 and N-2 of each residue form the repeating unit of the main chain, and the word "side chain" refers to the C-3 and higher numbered carbon atoms and their substituents. It also includes H atoms attached to the main chain atoms.

89

Amino acids may be classified according to the number of carbon atoms which appear in the main chain between the carbonyl carbon and amino nitrogen atoms which participate in the peptide bonds. Among the 150 or so amino acids which occur in nature, alpha, beta, gamma and delta amino acids are known. These have 1-4 intermediary carbons. Only alpha amino acids occur in proteins. Proline is a special case of an alpha amino acid; its side chain also binds to the peptide bond nitrogen.

For beta and higher order amino acids, there is a choice as to which main chain core carbon a side chain other than H is attached to. The preferred attachment site is the C-2 (alpha) carbon, i.e., the one adjacent to the carboxyl carbon of the -CO linking functionality. It is also possible for more than one main chain atom to carry a side chain other than H. However, in a preferred embodiment, only one main chain core atom carries a side chain other than H.

A main chain carbon atom may carry either one or two side chains; one is more common. A side chain may be attached to a main chain carbon atom by a single or a double bond; the former is more common.

A simple combinatorial peptide library is one whose members are peptides having three or more amino acids connected via peptide bonds.

The peptides may be linear, branched, or cyclic, and may covalently or noncovalently include nonpeptidyl moieties. The amino acids are not limited to the naturally occurring or to the genetically encoded amino acids.

A biased peptide library is one in which one or more (but not all) residues of the peptides are constant residues.

Cyclic Peptides

5

10

15

20

35

;0

٠5

Many naturally occurring peptides are cyclic.

Cyclization is a common mechanism for stabilization of peptide conformation thereby achieving improved association

90

of the peptide with its ligand and hence improved biological activity. Cyclization is usually achieved by intra-chain cystine formation, by formation of peptide bond between side chains or between N- and C- terminals. Cyclization was usually achieved by peptides in solution, but several publications have appeared that describe cyclization of peptides on beads.

A peptide library may be an oligopeptide library or a protein library.

Oligopeptides

5

LO

5

!lO

Preferably, the oligopeptides are at least five, six, seven or eight amino acids in length. Preferably, they are composed of less than 50, more preferably less than 20 amino acids.

In the case of an oligopeptide library, all or just some of the residues may be variable. The oligopeptide may be unconstrained, or constrained to a particular conformation by, e.g., the participation of constant cysteine residues in the formation of a constraining disulfide bond.

Proteins

Proteins, like oligopeptides, are composed of a plurality of amino acids, but the term protein is usually reserved for longer peptides, which are able to fold into a stable conformation. A protein may be composed of two or more polypeptide chains, held together by covalent or noncovalent crosslinks. These may occur in a homooligomeric or a heterooligomeric state.

A peptide is considered a protein if it (1) is at least 50 amino acids long, or (2) has at least two stabilizing covalent crosslinks (e.g., disulfide bonds). Thus, conotoxins are considered proteins.

Usually, the proteins of a protein library will be characterizable as having both constant residues (the same

91

for all proteins in the library) and variable residues (which vary from member to member). This is simply because, for a given range of variation at each position, the sequence space (simple diversity) grows exponentially with the number of residue positions, so at some point it becomes inconvenient for all residues of a peptide to be variable positions. Since proteins are usually larger than oligopeptides, it is more common for protein libraries than oligopeptide libraries to feature variable positions.

5

0

0

5

In the case of a protein library, it is desirable to focus the mutations at those sites which are tolerant of mutation. These may be determined by alanine scanning mutagenesis or by comparison of the protein sequence to that of homologous proteins of similar activity. It is also more likely that mutation of surface residues will directly affect binding. Surface residues may be determined by inspecting a 3D structure of the protein, or by labeling the surface and then ascertaining which residues have received labels. They may also be inferred by identifying regions of high hydrophilicity within the protein.

Because proteins are often altered at some sites but not others, protein libraries can be considered a special case of the biased peptide library.

There are several reasons that one might screen a protein library instead of an oligopeptide library, including (1) a particular protein, mutated in the library, has the desired activity to some degree already, and (2) the oligopeptides are not expected to have a sufficiently high affinity or specificity since they do not have a stable conformation.

When the protein library is based on a parental protein which does not have the desired activity, the parental protein will usually be one which is of high stability (melting point >= 50 deg. C.) and/or possessed of hypervariable regions.

The variable domains of an antibody possess

92

hypervariable regions and hence, in some embodiments, the protein library comprises members which comprise a mutant of VH or VL chain, or a mutant of an antigen-specific binding fragment of such a chain. VH and VL chains are usually each about 110 amino acid residues, and are held in proximity by a disulfide bond between the adjoing CL and CH1 regions to form a variable domain. Together, the VH, VL, CL and CH1 form an Fab fragment.

In human heavy chains, the hypervariable regions are at 31-35, 49-65, 98-111 and 84-88, but only the first three are involved in antigen binding. There is variation among VH and VL chains at residues outside the hypervariable regions, but to a much lesser degree.

A sequence is considered a mutant of a VH or VL chain if it is at least 80% identical to a naturally occurring VH or VL chain at all residues outside the hypervariable region.

In a preferred embodiment, such antibody library members comprise both at least one VH chain and at least one VL chain, at least one of which is a mutant chain, and which chains may be derived from the same or different antibodies. The VH and VL chains may be covalently joined by a suitable linker moiety, as in a "single chain antibody", or they may be noncovalently joined, as in a naturally occurring variable domain.

If the joining is noncovalent, and the library is displayed on cells or virus, then either the VH or the VL chain may be fused to the carrier surface/coat protein. The complementary chain may be co-expressed, or added exogenously to the library.

The members may further comprise some or all of an antibody constant heavy and/or constant light chain, or a mutant thereof.

Peptoid Library

5

)

A peptoid is an analogue of a peptide in which one or

93

more of the peptide bonds (-NH-CO-) are replaced by pseudopeptide bonds, which may be the same or different. It is not necessary that all of the peptide bonds be replaced, i.e., a peptoid may include one or more conventional amino acid residues, e.g., proline.

5

0

5

Э

5

A peptide bond has two small divalent linker elements, -NH- and -CO-. Thus, a preferred class of psuedopeptide bonds are those which consist of two small divalent linker elements. Each may be chosen independently from the group consisting of amine (-NH-), substituted amine (-NR-), carbonyl (-CO-), thiocarbonyl (-CS-), methylene (-CH2-), monosubstituted methylene (-CHR-), disubstituted methylene (-CR1R2-), ether (-O-) and thioether (-S-). The more preferred pseudopeptide bonds include:

N-modified -NRCOCarba Ψ -CH₂-CH₂Depsi Ψ -CO-OHydroxyethylene Ψ -CHOH-CH₂Ketomethylene Ψ -CO-CH₂Methylene-Oxy -CH₂-OReduced -CH₂-NHThiomethylene -CH₂-SThiopeptide -CS-NHRetro-Inverso -CO-NH-

A single peptoid molecule may include more than one kind of pseudopeptide bond.

For the purposes of introducing diversity into a peptoid library, one may vary (1) the side chains attached to the core main chain atoms of the monomers linked by the pseudopeptide bonds, and/or (2) the side chains (e.g., the -R of an -NRCO-) of the pseudopeptide bonds. Thus, in one embodiment, the monomeric units which are not amino acid residues are of the structure -NR1-CR2-CO-, where at least one of Rl and R2 are not hydrogen. If there is variability in the pseudopeptide bond, this is most conveniently done by

94

using an -NRCO- or other pseudopeptide bond with an R group, and varying the R group. In this event, the R group will usually be any of the side chains characterizing the amino acids of peptides, as previously discussed.

If the R group of the pseudopeptide bond is not variable, it will usually be small, e.g., not more than 10 atoms (e.g., hydroxyl, amino, carboxyl, methyl, ethyl, propyl).

If the conjugation chemistries are compatible, a simple combinatorial library may include both peptides and peptoids.

Peptide Nucleic Acid Library

5

.0

5

A PNA oligomer is here defined as one comprising a plurality of units, at least one of which is a PNA monomer which comprises a side chain comprising a nucleobase. For nucleobases, see USP 6,077,835.

The classic PNA oligomer is composed of (2-aminoethyl)glycine units, with nucleobases attached by methylene carbonyl linkers. That is, it has the structure

$$H - (-HN - CH_2 - CH_2 - N(-CO - CH_2 - B) - CH_2 - CO -)_n - OH$$

where the outer parenthesized substructure is the PNA monomer.

In this structure, the nucleobase B is separated from the backbone N by three bonds, and the points of attachment of the side chains are separated by six bonds. The nucleobase may be any of the bases included in the nucleotides discussed in connection with oligonucleotide libraries. The bases of nucleotides A, G, T, C and U are preferred.

A PNA oligomer may further comprise one or more amino acid residues, especially glycine and proline.

One can readily envision related molecules in which (1)

95

the -COCH2- linker is replaced by another linker, especially one composed of two small divalent linkers as defined previously, (2) a side chain is attached to one of the three main chain carbons not participating in the peptide bond (either instead or in addition to the side chain attached to the N of the classic PNA); and/or (3) the peptide bonds are replaced by pseudopeptide bonds as disclosed previously in the context of peptoids.

PNA oligomer libraries have been made; see e.g. Cook, 6,204,326.

Small Organic Compound Library

5

.0 -

.5

0

5

0

5

The small organic compound library ("compound library", for short) is a combinatorial library whose members are suitable for use as drugs if, indeed, they have the ability to mediate a biological activity of the target protein.

Peptides have certain disadvantages as drugs. These include susceptibility to degradation by serum proteases, and difficulty in penetrating cell membranes. Preferably, all or most of the compounds of the compound library avoid, or at least do not suffer to the same degree, one or more of the pharmaceutical disadvantages of peptides.

In designing a compound library, it is helpful to bear in mind the methods of molecular modification typically used to obtain new drugs. Three basic kinds of modification may be identified: disjunction, in which a lead drug is simplified to identify its component pharmacophoric moieties; conjunction, in which two or more known pharmacophoric moieties, which may be the same or different, are associated, covalently or noncovalently, to form a new drug; and alteration, in which one moiety is replaced by another which may be similar or different, but which is not in effect a disjunction or conjunction. The use of the terms "disjunction", "conjunction" and "alteration" is intended only to connote the structural relationship of the end product to the original leads, and not how the new drugs

96

are actually synthesized, although it is possible that the two are the same.

The process of disjunction is illustrated by the evolution of neostigmine (1931) and edrophonium (1952) from physostigmine (1925). Subsequent conjunction is illustrated by demecarium (1956) and ambenonium (1956).

5

.0

.5

Alterations may modify the size, polarity, or electron distribution of an original moiety. Alterations include ring closing or opening, formation of lower or higher homologues, introduction or saturation of double bonds, introduction of optically active centers, introduction, removal or replacement of bulky groups, isosteric or bioisosteric substitution, changes in the position or orientation of a group, introduction of alkylating groups, and introduction, removal or replacement of groups with a view toward inhibiting or promoting inductive (electrostatic) or conjugative (resonance) effects.

Thus, the substituents may include electron acceptors and/or electron donors. Typical electron donors (+I) include $-CH_3$, $-CH_2R$, $-CHR_2$, $-CR_3$ and $-COO^-$. Typical electron acceptors (-I) include $-NH_3+$, $-NR_3+$, $-NO_2$, -CN, -COOH, -COOR, -CHO, -COR, -COR,

The substituents may also include those which increase or decrease electronic density in conjugated systems. The former (+R) groups include -CH₃, -CR₃, -F, -C1, -Br, -I, -OH, -OR, -OCOR, -SH, -SR, -NH₂, -NR₂, and -NHCOR. The later (-R) groups include -NO₂, -CN, -CHC, -COR, -COOH, -COOR, -CONH₂, -SO₂R and -CF₃.

Synthetically speaking, the modifications may be achieved by a variety of unit processes, including nucleophilic and electrophilic substitution, reduction and oxidation, addition elimination, double bond cleavage, and cyclization.

For the purpose of constructing a library, a compound, or a family of compounds, having one or more pharmacological

97

activities (which need not be related to the known or suspected activities of the target protein), may be disjoined into two or more known or potential pharmacophoric moieties. Analogues of each of these moieties may be identified, and mixtures of these analogues reacted so as to reassemble compounds which have some similarity to the original lead compound. It is not necessary that all members of the library possess moieties analogous to all of the moieties of the lead compound.

5

10

15

3.0

:5

.0

5

The design of a library may be illustrated by the example of the benzodiazepines. Several benzodiazepine drugs, including chlordiazepoxide, diazepam and oxazepam, have been used as anti-anxiety drugs. Derivatives of benzodiazepines have widespread biological activities; derivatives have been reported to act not only as anxiolytics, but also as anticonvulsants; cholecystokinin (CCK) receptor subtype A or B, kappa opioid receptor, platelet activating factor, and HIV transactivator Tat antagonists, and GPIIbIIa, reverse transcriptase and ras farnesyltransferase inhibitors.

The benzodiazepine structure has been disjoined into a 2-aminobenzophenone, an amino acid, and an alkylating agent. See Bunin, et al., Proc. Nat. Acad. Sci. USA, 91:4708 (1994). Since only a few 2-aminobenzophenone derivatives are commercially available, it was later disjoined into 2-aminoarylstannane, an acid chloride, an amino acid, and an alkylating agent. Bunin, et al., Meth. Enzymol., 267:448 (1996). The arylstannane may be considered the core structure upon which the other moieties are substituted, or all four may be considered equals which are conjoined to make each library member:

A basic library synthesis plan and member structure is shown in Figure 1 of Fowlkes, et al., U.S. Serial No. 08/740,671, incorporated by reference in its entirety. The acid chloride building block introduces variability at the R^1 site. The R^2 site is introduced by the amino acid, and the

98

R³ site by the alkylating agent. The R⁴ site is inherent in the arylstannane. Bunin, et al. generated a 1, 4benzodiazepine library of 11,200 different derivatives prepared from 20 acid chlorides, 35 amino acids, and 16 alkylating agents. (No diversity was introduced at R4; this group was used to couple the molecule to a solid phase.) According to the Available Chemicals Directory (HDL Information Systems, San Leandro CA), over 300 acid chlorides, 80 Fmoc-protected amino acids and 800 alkylating agents were available for purchase (and more, of course. could be synthesized). The particular moieties used were chosen to maximize structural dispersion, while limiting the numbers to those conveniently synthesized in the wells of a microtiter plate. In choosing between structurally similar compounds, preference was given to the least substituted compound.

5

LO

.j5

5

0

5

The variable elements included both aliphatic and aromatic groups. Among the aliphatic groups, both acyclic and cyclic (mono- or poly-) structures, substituted or not, were tested. (While all of the acyclic groups were linear, it would have been feasible to introduce a branched aliphatic). The aromatic groups featured either single and multiple rings, fused or not, substituted or not, and with heteroatoms or not. The secondary substitutents included - NH2, -OH, -OMe, -CN, -C1, -F, and -COOH. While not used, spacer moieties, such as -O-, -S-, -OO-, -CS-, -NH-, and -NR-, could have been incorporated.

Bunin et al. suggest that instead of using a 1, 4-benzodiazepine as a core structure, one may instead use a 1, 4-benzodiazepine-2, 5-dione structure.

As noted by Bunin et al., it is advantageous, although not necessary, to use a linkage strategy which leaves no trace of the linking functionality, as this permits construction of a more diverse library.

Other combinatorial nonoligomeric compound libraries known or suggested in the art have been based on carbamates,

99

mercaptoacylated pyrrolidines, phenolic agents, aminimides, N-acylamino ethers (made from amino alcohols, aromatic hydroxy acids, and carboxylic acids), N-alkylamino ethers (made from aromatic hydroxy acids, amino alcohols and aldehydes) 1, 4-piperazines, and 1, 4-piperazine-6-ones.

5

LO

.5

:0

0

5

DeWitt, et al., Proc. Nat. Acad. Sci. (USA), 90:6909-13 (1993) describe the simultaneous but separate, synthesis of 40 discrete hydantoins and 40 discrete benzodiazepines. They carry out their synthesis on a solid support (inside a gas dispersion tube), in an array format, as opposed to other conventional simultaneous synthesis techniques (e.g., in a well, or on a pin). The hydantoins were synthesized by first simultaneously deprotecting and then treating each of five amino acid resins with each of eight isocyanates. The benzodiazepines were synthesized by treating each of five deprotected amino acid resins with each of eight 2-amino benzophenone imines.

Chen, et al., J. Am. Chem. Soc., 116:2661-62 (1994) described the preparation of a pilot (9 member) combinatorial library of formate esters. A polymer beadbound aldehyde preparation was "split" into three aliquots, each reacted with one of three different ylide reagents. The reaction products were combined, and then divided into three new aliquots, each of which was reacted with a different Michael donor. Compound identity was found to be determinable on a single bead basis by gas chromatography/mass spectroscopy analysis.

Holmes, USP 5,549,974 (1996) sets forth methodologies for the combinatorial synthesis of libraries of thiazolidinones and metathiazanones. These libraries are made by combination of amines, carbonyl compounds, and thiols under cyclization conditions.

Ellman, USP 5,545,568 (1996) describes combinatorial synthesis of benzodiazepines, prostaglandins, beta-turn mimetics, and glycerol-based compounds. See also Ellman, USP 5,288,514.

100

Summerton, USP 5,506,337 (1996) discloses methods of preparing a combinatorial library formed predominantly of morpholino subunit structures.

Heterocylic combinatorial libraries are reviewed generally in Nefzi, et al., Chem. Rev., 97:449-472 (1997).

5

0

For pharmacological classes, see, e.g., Goth, Medical Pharmacology: Principles and Concepts (C.V. Mosby Co.: 8th ed. 1976); Korolkovas and Burckhalter, Essentials of Medicinal Chemistry (John Wiley & Sons, Inc.: 1976). For synthetic methods, see, e.g., Warren, Organic Synthesis: The Disconnection Approach (John Wiley & Sons, Ltd.: 1982); Fuson, Reactions of Organic Compounds (John Wiley & Sons: 1966); Payne and Payne, How to do an Organic Synthesis (Allyn and Bacon, Inc.: 1969); Greene, Protective Groups in Organic Synthesis (Wiley-Interscience). For selection of substituents, see e.g., Hansch and Leo, Substituent Constants for Correlation Analysis in Chemistry and Biology (John Wiley & Sons: 1979).

The library is preferably synthesized so that the individual members remain identifiable so that, if a member is shown to be active, it is not necessary to analyze it. Several methods of identification have been proposed, including:

- (1) encoding, i.e., the attachment to each member of an identifier moiety which is more readily identified than the member proper. This has the disadvantage that the tag may itself influence the activity of the conjugate.
- (2) spatial addressing, e.g., each member is synthesized only at a particular coordinate on or in a matrix, or in a particular chamber. This might be, for example, the location of a particular pin, or a particular well on a microtiter plate, or inside a "tea bag".

The present invention is not limited to any particular form

101

of identification.

5

10

20

25

30

35

However, it is possible to simply characterize those members of the library which are found to be active, based on the characteristic spectroscopic indicia of the various building blocks.

Solid phase synthesis permits greater control over which derivatives are formed. However, the solid phase could interfere with activity. To overcome this problem, some or all of the molecules of each member could be liberated, after synthesis but before screening.

Examples of candidate simple libraries which might be evaluated include derivatives of the following:

Cyclic Compounds Containing One Hetero Atom

Heteronitrogen

15 pyrroles

pentasubstituted pyrroles

pyrrolidines

pyrrolines

prolines

indoles

beta-carbolines

pyridines

dihydropyridines

1,4-dihydropyridines

pyrido[2,3-d]pyrimidines

tetrahydro-3H-imidazo[4,5-c] pyridines

Isoquinolines

tetrahydroisoguinolines

quinolones

beta-lactams

azabicyclo[4.3.0]nonen-8-one amirno acid

Heterooxygen

furans

tetrahydrofurans

2,5-disubstituted tetrahydrofurans

pyrans 💰

102

hydroxypyranones

tetrahydroxypyranones

gamma-butyrolactones

Heterosulfur

sulfolenes

Cyclic Compounds with Two or More Hetero atoms

Multiple heteronitrogens

imidazoles

pyrazoles

piperazines

diketopiperazines

arylpiperazines

benzylpiperazines

benzodiazepines

1,4-benzodiazepine-2,5-diones

hydantoins

5-alkoxyhydantoins

dihydropyrimidines

1,3-disubstituted-5,6-dihydopyrimidine-2,4-

diones

cyclic ureas

cyclic thioureas

quinazolines

chiral 3-substituted-quinazoline-2,4-

diones

triazoles

1,2,3-triazoles

purines

Heteronitrogen and Heterooxygen

dikelomorpholines

isoxazoles

isoxazolinės

Heteronitrogen and Heterosulfur

thiazolidines

N-axylthiazolidines

103

dihydrothiazoles

2-methylene-2,3-dihydrothiazates

2-aminothiazoles

thiophenes

5

.0

5

0

5

5

3-amino thiophenes

4-thiazolidinones

4-melathiazanones

benzisothiazolones

For details on synthesis of libraries, see Nefzi, et al., Chem. Rev., 97:449-72 (1997), and references cited therein.

Pharmaceutical Methods and Preparations

The preferred animal subject of the present invention is a mammal. By the term "mammal" is meant an individual belonging to the class Mammalia. The invention is particularly useful in the treatment of human subjects, although it is intended for veterinary and nutritional uses as well. Preferred nonhuman subjects are of the orders Primata (e.g., apes and monkeys), Artiodactyla or Perissodactyla (e.g., cows, pigs, sheep, horses, goats), Carnivora (e.g., cats, dogs), Rodenta (e.g., rats, mice, guinea pigs, hamsters), Lagomorpha (e.g., rabbits) or other pet, farm or laboratory mammals.

The term "protection", as used herein, is intended to include "prevention," "suppression" and "treatment."
"Prevention", strictly speaking, involves administration of the pharmaceutical prior to the induction of the disease (or other adverse clinical condition). "Suppression" involves administration of the composition prior to the clinical appearance of the disease. "Treatment" involves administration of the protective composition after the appearance of the disease.

It will be understood that in human and veterinary medicine, it is not always possible to distinguish between "preventing" and "suppressing" since the ultimate inductive

104

event or events may be unknown, latent, or the patient is not ascertained until well after the occurrence of the event or events. Therefore, unless qualified, the term "prevention" will be understood to refer to both prevention in the strict sense, and to suppression.

The preventative or prophylactic use of a pharmaceutical usually involves identifying subjects who are at higher risk than the general population of contracting the disease, and administering the pharmaceutical to them in advance of the clinical appearance of the disease. The effectiveness of such use is measured by comparing the subsequent incidence or severity of the disease, or of particular symptoms of the disease, in the treated subjects against that in untreated subjects of the same high risk group.

While high risk factors vary from disease to disease, in general, these include (1) prior occurrence of the disease in one or more members of the same family, or, in the case of a contagious disease, in individuals with whom the subject has come into potentially contagious contact at a time when the earlier victim was likely to be contagious, (2) a prior occurrence of the disease in the subject, (3) prior occurrence of a related disease, or a condition known to increase the likelihood of the disease, in the subject; (4) appearance of a suspicious level of a marker of the disease, or a related disease or condition; (5) a subject who is immunologically compromised, e.g., by radiation treatment, HIV infection, drug use,, etc., or (6) membership in a particular group (e.g., a particular age, sex, race, ethnic group, etc.) which has been epidemiologically associated with that disease.

In some cases, it may be desirable to provide prophylaxis for the general population, and not just a high risk group. This is most likely to be the case when essentially all are at risk of contracting the disease, the effects of the disease are serious, the therapeutic index of

1.05

the prophylactic agent is high, and the cost of the agent is low.

A prophylaxis or treatment may be curative, that is, directed at the underlying cause of a disease, or ameliorative, that is, directed at the symptoms of the disease, especially those which reduce the quality of life.

5

0

5

0

5

5

It should also be understood that to be useful, the protection provided need not be absolute, provided that it is sufficient to carry clinical value. An agent which provides protection to a lesser degree than do competitive agents may still be of value if the other agents are ineffective for a particular individual, if it can be used in combination with other agents to enhance the level of protection, or if it is safer than competitive agents. It is desirable that there be a statistically significant (p=0.05 or less) improvement in the treated subject relative to an appropriate untreated control, and it is desirable that this improvement be at least 10%, more preferably at least 25%, still more preferably at least 50%, even more preferably at least 100%, in some indicia of the incidence or severity of the disease or of at least one symptom of the disease.

At least one of the drugs of the present invention may be administered, by any means that achieve their intended purpose, to protect a subject against a disease or other adverse condition. The form of administration may be systemic or topical. For example, administration of such a composition may be by various parenteral routes such as subcutaneous, intravenous, intradermal, intramuscular, intraperitoneal, intranasal, transdermal, or buccal routes. Alternatively, or concurrently, administration may be by the oral route. Parenteral administration can be by bolus injection or by gradual perfusion over time.

A typical regimen comprises administration of an effective amount of the drug, administered over a period ranging from a single dose, to dosing over a period of hours, days, weeks, months, or years.

106

It is understood that the suitable dosage of a drug of the present invention will be dependent upon the age, sex, health, and weight of the recipient, kind of concurrent treatment, if any, frequency of treatment, and the nature of the effect desired. However, the most preferred dosage can be tailored to the individual subject, as is understood and determinable by one of skill in the art, without undue experimentation. This will typically involve adjustment of a standard dose, e.g., reduction of the dose if the patient has a low body weight.

5

10

15

20

35

3 Ó

35

Prior to use in humans, a drug will first be evaluated for safety and efficacy in laboratory animals. In human clinical studies, one would begin with a dose expected to be safe in humans, based on the preclinical data for the drug in question, and on customary doses for amalogous drugs (if If this dose is effective, the dosage may be decreased, to determine the minimum effective dose, if desired. If this dose is ineffective, it will be cautiously increased, with the patients monitored for signs of side effects. See, e.g., Berkow et al, eds., The Merck Manual, 15th edition, Merck and Co., Rahway, N.J., 1987; Goodman et al., eds., Goodman and Gilman's The Pharmacological Basis of Therapeutics, 8th edition, Pergamon Press, Inc., Elmsford, N.Y., (1990); Avery's Drug Treatment: Principles and Practice of Clinical Pharmacology and Therapeutics, 3rd edition, ADIS Press, LTD., Williams and Wilkins, Baltimore, MD. (1987), Ebadi, Pharmacology, Little, Brown and Co., Boston, (1985), which references and references cited therein, are entirely incorporated herein by reference.

The total dose required for each treatment may be administered by multiple doses or in a single dose. The protein may be administered alone or in conjunction with other therapeutics directed to the disease or directed to other symptoms thereof.

Typical pharmaceutical doses, for adult humans, are in the range of 1 ng to 10g per day, more often 1 mg to 1g per

107

day.

5

10

15

20

25

30

;5

The appropriate dosage form will depend on the disease, the pharmaceutical, and the mode of administration; possibilities include tablets, capsules, lozenges, dental pastes, suppositories, inhalants, solutions, ointments and parenteral depots. See, e.g., Berker, supra, Goodman, supra, Avery, supra and Ebadi, supra, which are entirely incorporated herein by reference, including all references cited therein.

In the case of peptide drugs, the drug may be administered in the form of an expression vector comprising a nucleic acid encoding the peptide; such a vector, after incorporation into the genetic complement of a cell of the patient, directs synthesis of the peptide. Suitable vectors include genetically engineered poxviruses (vaccinia), adenoviruses, adeno-associated viruses, herpesviruses and lentiviruses which are or have been rendered nonpathogenic.

In addition to at least one drug as described herein, a pharmaceutical composition may contain suitable pharmaceutically acceptable carriers, such as excipients, carriers and/or auxiliaries which facilitate processing of the active compounds into preparations which can be used pharmaceutically. See, e.g., Berker, supra, Goodman, supra, Avery, supra and Ebadi, supra, which are entirely incorporated herein by reference, included all references cited therein.

Assay Compositions and Methods

Target Organism

The invention contemplates that it may be appropriate to ascertain or to mediate the biological activity of a substance of this invention in a target organism.

The target organism may be a plant, animal, or microorganism.

In the case of a plant, it may be an economic plant, in which case the drug may be intended to increase the disease,

108

weather or pest resistance, alter the growth characteristics, or otherwise improve the useful characteristics or mute undesirable characteristics of the plant. Or it may be a weed, in which case the drug may be intended to kill or otherwise inhibit the growth of the plant, or to alter its characteristics to convert it from a weed to an economic plant. The plant may be a tree, shrub, crop, grass, etc. The plant may be an algae (which are in some cases also microorganisms), or a vascular plant, especially gymnosperms (particularly conifers) and angiosperms. Angiosperms may be monocots or dicots. The plants of greatest interest are rice, wheat, corn, alfalfa, soybeans, potatoes, peanuts, tomatoes, melons, apples, pears, plums, pineapples, fir, spruce, pine, cedar, and oak.

If the target organism is a microorganism, it may be algae, bacteria, fungi, or a virus (although the biological activity of a virus must be determined in a virus-infected cell). The microorganism may be human or other animal or plant pathogen, or it may be nonpathogenic. It may be a soil or water organism, or one which normally lives inside other living things.

If the target organism is an animal, it may be a vertebrate or a nonvertebrate animal. Nonvertebrate animals are chiefly of interest when they act as pathogens or parasites, and the drugs are intended to act as biocidic or biostatic agents. Nonvertebrate animals of interest include worms, mollusks, and arthropods.

The target organism may also be a vertebrate animal, i.e., a mammal, bird, reptile, fish or amphibian. Among mammals, the target animal preferably belongs to the order Primata (humans, apes and monkeys), Artiodactyla (e.g., cows, pigs, sheep, goats, horses), Rodenta (e.g., mice, rats) Lagomorpha (e.g., rabbits, hares), or Carnivora (e.g., cats, dogs). Among birds, the target animals are preferably of the orders Anseriformes (e.g., ducks, geese, swans) or Galliformes (e.g., quails, grouse, pheasants, turkeys and

109

chickens). Among fish, the target animal is preferably of the order Clupeiformes (e.g., sardines, shad, anchovies, whitefish, salmon).

<u>Tarqet Tissues</u>

5

0

5

)

The term "target tissue" refers to any whole animal, physiological system, whole organ, part of organ, miscellaneous tissue, cell, or cell component (e.g., the cell membrane) of a target animal in which biological activity may be measured.

Routinely in mammals one would choose to compare and contrast the biological impact on virtually any and all tissues which express the subject receptor protein. The main tissues to use are: brain, heart, lung, kidney, liver, pancreas, skin, intestines, adipose, stomach, skeletal muscle, adrenal glands, breast, prostate, vasculature, retina, cornea, thyroid gland, parathyroid glands, thymus, bone marrow, bone, etc.

Another classification would be by cell type: B cells, T cells, macrophages, neutrophils, eosinophils, mast cells, platelets, megakaryocytes, erythrocytes, bone marrow stomal cells, fibroblasts, neurons, astrocytes, neuroglia, microglia, epithelial cells (from any organ, e.g. skin, breast, prostate, lung, intestines etc), cardiac muscle cells, smooth muscle cells, striated muscle cells, osteoblasts, osteocytes, chondroblasts, chondrocytes, keratinocytes, melanocytes, etc.

Of course, in the case of a unicellular organism, there is no distinction between the "target organism" and the "target tissue".

Screening Assays

Assays intended to determine the binding or the biological activity of a substance are called preliminary screening assays.

Screening assays will typically be either in vitro

110

(cell-free) assays (for binding to an immobilized receptor) or cell-based assays (for alterations in the phenotype of the cell). They will not involve screening of whole multicellular organisms, or isolated organs. The comments on diagnostic biological assays apply mutatis mutandis to screening cell-based assays.

In Vitro vs. In Vivo Assays

5

0

5

The term in vivo is descriptive of an event, such as binding or enzymatic action, which occurs within a living organism. The organism in question may, however, be genetically modified. The term in vitro refers to an event which occurs outside a living organism. Parts of an organism (e.g., a membrane, or an isolated biochemical) are used, together with artificial substrates and/or conditions. For the purpose of the present invention, the term in vitro excludes events occurring inside or on an intact cell, whether of a unicellular or multicellular organism.

In vivo assays include both cell-based assays, and organismic assays. The cell-based assays include both assays on unicellular organisms, and assays on isolated cells or cell cultures derived from multicellular organisms. The cell cultures may be mixed, provided that they are not organized into tissues or organs. The term organismic assay refers to assays on whole multicellular organisms, and assays on isolated organs or tissues of such organisms.

In vitro Diagnostic Methods and Reagents

The in vitro assays of the present invention may be applied to any suitable analyte-containing sample, and may be qualitative or quantitative in nature.

Sample

5

The sample will normally be a biological fluid, such as blood, urine, lymph, semen, milk, or cerebrospinal fluid, or

111

a fraction or derivative thereof, or a biological tissue, in the form of, e.g., a tissue section or homogenate. However, the sample conceivably could be (or derived from) a food or beverage, a pharmaceutical or diagnostic composition, soil, or surface or ground water. If a biological fluid or tissue, it may be taken from a human or other mammal, vertebrate or animal, or from a plant. The preferred sample is blood, or a fraction or derivative thereof.

Binding and Reaction Assays

0

) _

The assay may be a binding assay, in which one step involves the binding of a diagnostic reagent to the analyte, or a reaction assay, which involves the reaction of a reagent with the analyte. The reagents used in a binding assay may be classified as to the nature of their interaction with analyte: (1) analyte analogues, or (2) analyte binding molecules (ABM). They may be labeled or insolubilized.

In a reaction assay, the assay may look for a direct reaction between the analyte and a reagent which is reactive with the analyte, or if the analyte is an enzyme or enzyme inhibitor, for a reaction catalyzed or inhibited by the analyte. The reagent may be a reactant, a catalyst, or an inhibitor for the reaction.

An assay may involve a cascade of steps in which the product of one step acts as the target for the next step. These steps may be binding steps, reaction steps, or a combination thereof.

Signal Producing System (SPS)

In order to detect the presence, or measure the amount, of an analyte, the assay must provide for a signal producing system (SPS) in which there is a detectable difference in the signal produced, depending on whether the analyte is present or absent (or, in a quantitative assay, on the

112

amount of the analyte). The detectable signal may be one which is visually detectable, or one detectable only with instruments. Possible signals include production of colored or luminescent products, alteration of the characteristics (including amplitude or polarization) of absorption or emission of radiation by an assay component or product, and precipitation or agglutination of a component or product. The term "signal" is intended to include the discontinuance of an existing signal, or a change in the rate of change of an observable parameter, rather than a change in its absolute value. The signal may be monitored manually or automatically.

In a reaction assay, the signal is often a product of the reaction. In a binding assay, it is normally provided by a label borne by a labeled reagent.

Labels

10

Ľ5

30

:5

The component of the signal producing system which is most intimately associated with the diagnostic reagent is called the "label". A label may be, e.g., a radioisotope, a fluorophore, an enzyme, a co-enzyme, an enzyme substrate, an electron-dense compound, an agglutinable particle.

The radioactive isotope can be detected by such means as the use of a gamma counter or a scintillation counter or by autoradiography. Isotopes which are particularly useful for the purpose of the present invention include ³H, ¹²⁵I, ¹³¹I, ³⁵S, ¹⁴C, ³²P and ³³P. ¹²⁵I is preferred for antibody labeling.

The label may also be a fluorophore. When the fluorescently labeled reagent is exposed to light of the proper wave length, its presence can then be detected due to fluorescence. Among the most commonly used fluorescent labeling compounds are fluorescein isothiocyanate, rhodamine, phycoerythrin, phycocyanin, allophycocyanin, ophthaldehyde and fluorescamine.

Alternatively, fluorescence-emitting metals such as

113

125 Eu, or others of the lanthanide series, may be incorporated into a diagnostic reagent using such metal chelating groups as diethylenetriaminepentaacetic acid (DTPA) of ethylenediamine-tetraacetic acid (EDTA).

The label may also be a chemiluminescent compound. The presence of the chemiluminescently labeled reagent is then determined by detecting the presence of luminescence that arises during the course of a chemical reaction. Examples of particularly useful chemiluminescent labeling compounds are luminol, isolumino, theromatic acridinium ester, imidazole, acridinium salt and oxalate ester.

Likewise, a bioluminescent compound may be used for labeling. Bioluminescence is a type of chemiluminescence found in biological systems in which a catalytic protein increases the efficiency of the chemiluminescent reaction. The presence of a bioluminescent protein is determined by detecting the presence of luminescence. Important bioluminescent compounds for purposes of labeling are luciferin, luciferase and aequorin.

Enzyme labels, such as horseradish peroxidase and alkaline phosphatase, are preferred. When an enzyme label is used, the signal producing system must also include a substrate for the enzyme. If the enzymatic reaction product is not itself detectable, the SPS will include one or more additional reactants so that a detectable product appears.

An enzyme analyte may act as its own label if an enzyme inhibitor is used as a diagnostic reagent.

Binding Assay Formats

5

.0

5

0

5

0

5

Binding assays may be divided into two basic types, heterogeneous and homogeneous. In heterogeneous assays, the interaction between the affinity molecule and the analyte does not affect the label, hence, to determine the amount or presence of analyte, bound label must be separated from free label. In homogeneous assays, the interaction does affect the activity of the label, and therefore analyte levels can

114

be deduced without the need for a separation step.

5

10

15

30

:5

0

5

In one embodiment, the ABM is insolubilized by coupling it to a macromolecular support, and analyte in the sample is allowed to compete with a known quantity of a labeled or specifically labelable analyte analogue. The "analyte analogue" is a molecule capable of competing with analyte for binding to the ABM, and the term is intended to include analyte itself. It may be labeled already, or it may be labeled subsequently by specifically binding the label to a moiety differentiating the analyte analogue from analyte. The solid and liquid phases are separated, and the labeled analyte analogue in one phase is quantified. The higher the level of analyte analogue in the solid phase, i.e., sticking to the ABM, the lower the level of analyte in the sample.

In a "sandwich assay", both an insolubilized ABM, and a labeled ABM are employed. The analyte is captured by the insolubilized ABM and is tagged by the labeled ABM, forming a ternary complex. The reagents may be added to the sample in either order, or simultaneously. The ABMs may be the same or different. The amount of labeled ABM in the ternary complex is directly proportional to the amount of analyte in the sample.

The two embodiments described above are both heterogeneous assays. However, homogeneous assays are conceivable. The key is that the label be affected by whether or not the complex is formed. Conjugation Methods

A label may be conjugated, directly or indirectly (e.g., through a labeled anti-ABM antibody), covalently (e.g., with SPDP) or noncovalently, to the ABM, to produce a diagnostic reagent. Similarly, the ABM may be conjugated to a solid phase support to form a solid phase ("capture") diagnostic reagent.

Suitable supports include glass, polystyrene, polypropylene, polyethylene, dextran, nylon, amylases,

115

natural and modified celluloses, polyacrylamides, agaroses, and magnetite. The nature of the carrier can be either soluble to some extent or insoluble for the purposes of the present invention.

The support material may have virtually any possible structural configuration so long as the coupled molecule is capable of binding to its target. Thus the support configuration may be spherical, as in a bead, or cylindrical, as in the inside surface of a test tube, or the external surface of a rod. Alternatively, the surface may be flat such as a sheet, test strip, etc.

Biological Assays

5 .

.0

.5

0

5

0

5

A biological assay measures or detects a biological response of a biological entity to a substance.

The biological entity may be a whole organism, an isolated organ or tissue, freshly isolated cells, an immortalized cell line, or a subcellular component (such as a membrane; this term should not be construed as including an isolated receptor). The entity may be, or may be derived from, an organism which occurs in nature, or which is modified in some way. Modifications may be genetic (including radiation and chemical mutants, and genetic engineering) or somatic (e.g., surgical, chemical, etc.). In the case of a multicellular entity, the modifications may affect some or all cells. The entity need not be the target organism, or a derivative thereof, if there is a reasonable correlation between bioassay activity in the assay entity and biological activity in the target organism.

The entity is placed in a particular environment, which may be more or less natural. For example, a culture medium may, but need not, contain serum or serum substitutes, and it may, but need not, include a support matrix of some kind, it may be still, or agitated. It may contain particular biological or chemical agents, or have particular physical parameters (e.g., temperature), that are intended to nourish

116

or challenge the biological entity.

10

Ĺ5

10

:5

Q

There must also be a detectable biological marker for the response. At the cellular level, the most common markers are cell survival and proliferation, cell behavior (clustering, motility), cell morphology (shape, color), and biochemical activity (overall DNA synthesis, overall protein synthesis, and specific metabolic activities, such as utilization of particular nutrients, e.g., consumption of oxygen, production of CO_2 , production of organic acids, uptake or discharge of ions).

The direct signal produced by the biological marker may be transformed by a signal producing system into a different signal which is more observable, for example, a fluorescent or colorimetric signal.

The entity, environment, marker and signal producing system are chosen to achieve a clinically acceptable level of sensitivity, specificity and accuracy.

In some cases, the goal will be to identify substances which mediate the biological activity of a natural biological entity, and the assay is carried out directly with that entity. In other cases, the biological entity is used simply as a model of some more complex (or otherwise inconvenient to work with) biological entity. In that event, the model biological entity is used because activity in the model system is considered more predictive of activity in the ultimate natural biological entity than is simple binding activity in an in vitro system. The model entity is used instead of the ultimate entity because the former is more expensive or slower to work with, or because ethical considerations forbid working with the ultimate entity yet.

The model entity may be naturally occurring, if the model entity usefully models the ultimate entity under some conditions. Or it may be non-naturally occurring, with modifications that increase its resemblance to the ultimate entity.

117

Transgenic animals, such as transgenic mice, rats, and rabbits, have been found useful as model systems.

In cell-based model assays, where the biological activity is mediated by binding to a receptor (target protein), the receptor may be functionally connected to a signal (biological marker) producing system, which may be endogenous or exogenous to the cell.

There are a number of techniques of doing this.

"Zero-Hybrid" Systems

5

LO.

.5

10

:5

0

5

In these systems, the binding of a peptide to the target protein results in a screenable or selectable phenotypic change, without resort to fusing the target protein (or a ligand binding moiety thereof) to an endogenous protein. It may be that the target protein is endogenous to the host cell, or is substantially identical to an endogenous receptor so that it can take advantage of the latter's native signal transduction pathway. Or sufficient elements of the signal transduction pathway normally associated with the target protein may be engineered into the cell so that the cell signals binding to the target protein.

"One-Hybrid" Systems

In these systems, a chimera receptor, a hybrid of the target protein and an endogenous receptor, is used. The chimeric receptor has the ligand binding characteristics of the target protein and the signal transduction characteristics of the endogenous receptor. Thus, the normal signal transduction pathway of the endogenous receptor is subverted.

Preferably, the endogenous receptor is inactivated, or the conditions of the assay avoid activation of the endogenous receptor, to improve the signal-to-noise ratio.

See Fowlkes USP 5,789,184 for a yeast system.

Another type of "one-hybrid" system combines a peptide:

1.18

DNA-binding domain fusion with an unfused target receptor that possesses an activation domain.

"Two-Hybrid" System

5

ĹΟ

15

0.5

· 0,

.5

In a preferred embodiment, the cell-based assay is a two hybrid system. This term implies that the ligand is incorporated into a first hybrid protein, and the receptor into a second hybrid protein. The first hybrid also comprises component A of a signal generating system, and the second hybrid comprises component B of that system.

Components A and B, by themselves, are insufficient to generate a signal. However, if the ligand binds the receptor, components A and B are brought into sufficiently close proximity so that they can cooperate to generate a signal.

Components A and B may naturally occur, or be substantially identical to moieties which naturally occur, as components of a single naturally occurring biomolecule, or they may naturally occur, or be substantially identical to moieties which naturally occur, as separate naturally occurring biomolecules which interact in nature.

Two-Hybrid System: Transcription Factor Type

In a preferred "two-hybrid" embodiment, one member of a peptide ligand:receptor binding pair is expressed as a fusion to a DNA-binding domain (DBD) from a transcription factor (this fusion protein is called the "bait"), and the other is expressed as a fusion to a transactivation domain (TAD) (this fusion protein is called the "fish", the "prey", or the "catch"). The transactivation domain should be complementary to the DNA-binding domain, i.e., it should interact with the latter so as to activate transcription of a specially designed reporter gene that carries a binding site for the DNA-binding domain. Naturally, the two fusion proteins must likewise be complementary.

This complementarity may be achieved by use of the

119

complementary and separable DNA-binding and transcriptional activator domains of a single transcriptional activator protein, or one may use complementary domains derived from different proteins. The domains may be identical to the native domains, or mutants thereof. The assay members may be fused directly to the DBD or TAD, or fused through an intermediated linker.

5

٥.

.5

:0

5

0

5

The target DNA operator may be the native operator sequence, or a mutant operator. Mutations in the operator may be coordinated with mutations in the DBD and the TAD. An example of a suitable transcription activation system is one comprising the DNA-binding domain from the bacterial repressor LexA and the activation domain from the yeast transcription factor Gal4, with the reporter gene operably linked to the LexA operator.

It is not necessary to employ the intact target receptor; just the ligand-binding moiety is sufficient.

The two fusion proteins may be expressed from the same or different vectors. Likewise, the activatable reporter gene may be expressed from the same vector as either fusion protein (or both proteins), or from a third vector.

Potential DNA-binding domains include Gal4, LexA, and mutant domains substantially identical to the above.

Potential activation domains include E. coli B42, Gal4 activation domain II, and HSV VP16, and mutant domains substantially identical to the above.

Potential operators include the native operators for the desired activation domain, and mutant domains substantially identical to the native operator.

The fusion proteins may comprise nuclear localization signals.

The assay system will include a signal producing system, too. The first element of this system is a reporter gene operably linked to an operator responsive to the DBD and TAD of choice. The expression of this reporter gene will result, directly or indirectly, in a selectable or

120

screenable phenotype (the signal). The signal producing system may include, besides the reporter gene, additional genetic or biochemical elements which cooperate in the production of the signal. Such an element could be, for example, a selective agent in the cell growth medium. There may be more than one signal producing system, and the system may include more than one reporter gene.

The sensitivity of the system may be adjusted by, e.g., use of competitive inhibitors of any step in the activation or signal production process, increasing or decreasing the number of operators, using a stronger or weaker DBD or TAD, etc.

When the signal is the death or survival of the cell in question, or proliferation or nonproliferation of the cell in question, the assay is said to be a selection. When the signal merely results in a detectable phenotype by which the signaling cell may be differentiated from the same cell in a nonsignaling state (either way being a living cell), the assay is a screen. However, the term "screening assay" may be used in a broader sense to include a selection. When the narrower sense is intended, we will use the term "nonselective screen".

.5

:O

Various screening and selection systems are discussed in Ladner, USP 5,198,346.

Screening and selection may be for or against the peptide: target protein or compound:target protein interaction.

Preferred assay cells are microbial (bacterial, yeast, algal, protozooal), invertebrate, vertebrate (esp. mammalian, particularly human). The best developed two-hybrid assays are yeast and mammalian systems.

Normally, two hybrid assays are used to determine whether a protein X and a protein Y interact, by virtue of their ability to reconstitute the interaction of the DBD and the TAD. However, augmented two-hybrid assays have been used to detect interactions that depend on a third, non-

121

protein ligand.

5

10

15

20

25

30

35

For more guidance on two-hybrid assays, see Brent and Finley, Jr., Ann. Rev. Genet., 31:663-704 (1997); Fremont-Racine, et al., Nature Genetics, 277-281 (16 July 1997); Allen, et al., TIBS, 511-16 (Dec. 1995); LeCrenier, et al., BioEssays, 20:1-6 (1998); Xu, et al., Proc. Nat. Acad. sci. (USA), 94:12473-8 (Nov. 1992); Esotak, et al., Mol. Cell. Biol., 15:5820-9 (1995); Yang, et al., Nucleic Acids Res., 23:1152-6 (1995); Bendixen, et al., Nucleic Acids Res., 22:1778-9 (1994); Fuller, et al., BioTechniques, 25:85-92 (July 1998); Cohen, et al., PNAS (USA) 95:14272-7 (1998); Kolonin and Finley, Jr., PNAS (USA) 95:14266-71 (1998). See also Vasavada, et al., PNAS (USA), 88:10686-90 (1991) (contingent replication assay), and Rehrauer, et al., J. Biol. Chem., 271:23865-73 91996) (LexA repressor cleavage assay).

Two-Hybrid Systems: reporter Enzyme type

In another embodiment, the components A and B reconstitute an enzyme which is not a transcription factor.

As in the last example, the effect of the reconstitution of the enzyme is a phenotypic change which may be a screenable change, a selectable change, or both.

In vivo Diagnostic Uses

Radio-labeled ABM may be administered to the human or animal subject. Administration is typically by injection, e.g., intravenous or arterial or other means of administration in a quantity sufficient to permit subsequent dynamic and/or static imaging using suitable radio-detecting devices. The dosage is the smallest amount capable of providing a diagnostically effective image, and may be determined by means conventional in the art, using known radio-imaging agents as a guide.

Typically, the imaging is carried out on the whole body

122

of the subject, or on that portion of the body or organ relevant to the condition or disease under study. The amount of radio-labeled ABM accumulated at a given point in time in relevant target organs can then be quantified.

A particularly suitable radio-detecting device is a scintillation camera, such as a gamma camera. scintillation camera is a stationary device that can be used to image distribution of radio-labeled ABM. The detection device in the camera senses the radioactive decay, 'the distribution of which can be recorded. Data produced by the imaging system can be digitized. The digitized information can be analyzed over time discontinuously or continuously. The digitized data can be processed to produce images, called frames, of the pattern of uptake of the radio-labeled ABM in the target organ at a discrete point in time. most continuous (dynamic) studies, quantitative data is obtained by observing changes in distributions of radioactive decay in target organs over time. In other words, a time-activity analysis of the data will illustrate uptake through clearance of the radio-labeled binding protein by the target organs with time.

Various factors should be taken into consideration in selecting an appropriate radioisotope. The radioisotope must be selected with a view to obtaining good quality resolution upon imaging, should be safe for diagnostic use in humans and animals, and should preferably have a short physical half-life so as to decrease the amount of radiation received by the body. The radioisotope used should preferably be pharmacologically inert, and, in the quantities administered, should not have any substantial physiological effect.

51

)

The ABM may be radio-labeled with different isotopes of iodine, for example ¹²³I, ¹²⁵I, or ¹³¹I (see for example, U.S. Patent 4,609,725). The extent of radio-labeling must, however be monitored, since it will affect the calculations made based on the imaging results (i.e. a diiodinated ABM

123

will result in twice the radiation count of a similar monoiodinated ABM over the same time frame).

0

0

5

Э

In applications to human subjects, it may be desirable to use radioisotopes other than ¹²⁵I for labeling in order to decrease the total dosimetry exposure of the human body and to optimize the detectability of the labeled molecule (though this radioisotope can be used if circumstances require). Ready availability for clinical use is also a factor. Accordingly, for human applications, preferred radio-labels are for example, ^{99m}Tc, ⁶⁷Ga, ⁶⁸Ga, ⁹⁰Y, ¹¹¹In, ^{113m}In, ¹²³I, ¹⁸⁶Re, ¹⁸⁸Re or ²¹¹At.

The radio-labeled ABM may be prepared by various methods. These include radio-halogenation by the chloramine - T method or the lactoperoxidase method and subsequent purification by HPLC (high pressure liquid chromatography), for example as described by J. Gutkowska et al in "Endocrinology and Metabolism Clinics of America: (1987) 16 (1):183. Other known methods of radio-labeling can be used, such as IODOBEADS™.

There are a number of different methods of delivering the radio-labeled ABM to the end-user. It may be administered by any means that enables the active agent to reach the agent's site of action in the body of a mammal. Because proteins are subject to being digested when administered orally, parenteral administration, i.e., intravenous, subcutaneous, intramuscular, would ordinarily be used to optimize absorption of an ABM, such as an antibody, which is a protein.

EXAMPLES

Example 1

Differentially expressed mouse genes, and corresponding human genes/proteins, were identified as described in this Example, and compiled into Master Table 1.

Animal Models Upon separation from their mothers (weaning), C57Bl/6J mice (i.e., C57Bl/6 mice developed by Jackson Labs) were placed on a normal diet (PMI Nutrition International Inc., Brentwood, MO, Prolab RMH3000). Mice were sacrificed at an average of 35, 49, 56, 77, 118, 133, 207, 403, 558 and 725 days of age.

RNA isolation.

Total RNA was isolated from livers using the RNA STAT-60 Total RNA/mRNA Isolation Reagent according to the manufacturer's instructions (Tel-Test, Friendswood, TX).

Sample Quantification and Quality Assessment

Total RNA was quantified and assessed for quality on a Bioanalyzer RNA 6000 Nano chip (Agilent). Each chip contained an interconnected set of gel-filled channels that allowed for molecular sieving of nucleic acids. Pinelectrodes in the chip were used to create electrokinetic forces capable of driving molecules through these microchannels to perform electrophoretic separations. Ribosomal peaks were measured by fluorescence signal and displayed in an electropherogram. A successful total RNA sample featured 2 distinct ribosomal peaks (18S and 28S rRNA).

Biotinylated cRNA Hybridization Target.

Total RNA was prepared for use as a hybridization target as described in the manufacturer's instructions for CodeLink Expression Bioarrays(TM) (Amersham Biosciences). The CodeLink Expression Bioarrays utilize nucleic acid

125

hybridization of a biotin-labeled complementary RNA(cRNA) target with DNA oligonucleotide probes attached to a gel matrix.

The biotin-labeled cRNA target is prepared by a linear amplification method. Poly (A) + RNA (within the total RNA population) is primed for reverse transcription by a DNA oligonucleotide containing a T7 RNA polymerase promoter 5' to a (dT) 24 sequence. After second-strand cDNA synthesis, the cDNA serves as the template in an *in vitro* transcription (IVT) reaction to produce the target cRNA. The IVT is performed in the presence of biotinylated nucleotides to label the target cRNA. This procedure results in a 50-200 fold linear amplification of the input poly (A) + RNA.

Hybridization Probes.

5

0

5

0

5

0

5

The oligonucleotide probes were provided by the Codelink Uniset Mouse I Bioarray (Amersham, product code 300013). Amine-terminated oligonucleotide probes are attached to a three-dimensional polyacrylamide gel matrix. There are 10,000 oligonucleotide probes, each specific to a well-characterized mouse gene. Each mouse gene is representative of a unique gene cluster from the fourth quarter 2001 Genbank Unigene build. There are also 500 control probes.

The sequences of the probes are proprietary to Amersham. However, for each probe, Amersham identifies the corresponding mouse gene by NCBI accession number, OGS, LocusLink, Unigene Cluster ID, and description (name). This information should be available from Amersham. In the case of the differentially expressed probes, this information is duplicated in master table 1. For the complete list, see http://www4.amershambiosciences.com/aptrix/upp01077.nsf/Content/codelink literature

Under "Gene Lists", select "Uniset Mouse I", and a gene

126

list, in Excel format, can be downloaded.

Hybridization

15

20

:5

Using the cRNA target, the hybridization reaction mixture is prepared and loaded into array chambers for bioarray processing as set forth in the manufacturer's instructions for CodeLink Gene Expression BioarraysTM (Amerhsam Biosciences). Each sample is hybridized to an individual microarray. Hybridization is at 37°C. The hybridization buffer is prepared as set forth in the Motorola instructions. Hybridization to the microarray is detected with an avidinated fluorescent reagent, Streptavidin-Alexa Fluor ® 647 (Amersham).

Mouse Gene Expression Analysis

Processed arrays were scanned using a GenePix 4000B Microarray Scanner (Axon Instruments, Inc.); array images were acquired using the Amersham CodeLink™ Analysis Software (Release 2.2). The Amersham CodeLink™ Analysis Software gives an integrated optical density (IOD) value for every spot; a unique background value for that spot is subtracted, resulting in "raw" data points. Individual chips are then normalized by the Amersham Codelink™ software according to the median raw intensity for all 10,000 genes. A negative control threshold (0.2) was also calculated according to the control probes. A significant difference in expression between samples was defined as a minimum of 2-fold change in expression values. Genes with expression values below the negative control threshold were eliminated from the analysis and then the expression data was analyzed to identify genes whose expression levels changed significantly with respect to age.

The list of genes in the tables is a combination of two analyses. Samples of average age 35, 49, 77 and 133 days were compared pair-wise in all possible combinations (6 comparisons) and genes showing differences in expression

127

greater than 2-fold were listed in the table. (The 56 day data was not included in the comparisons.) The remaining samples were divided into three groups (118 days (2 mice): young; 207 and 403 (4 mice) averaged together: medium; 558 and 725 (4 mice) averaged together: old), the three groups were compared in all possible pair-wise combinations (3 comparisons) and genes showing differences in expression greater than 2-fold were added to the table.

Database Searches Nucleotide sequences and predicted amino acid sequences were compared to public domain databases using the Blast 2.0 program (National Center for Biotechnology Information, National Institutes of Health). Nucleotide sequences were displayed using ABI prism Edit View 1.0.1 (PE Applied Biosystems, Foster City, CA).

Nucleotide database searches were conducted with the

128

never submitted to an archival database but is available in the literature. A small number of sequences are provided through collaboration; the underlying primary sequence data is available in GenBank, but may not be available in any one GenBank record. RefSeq sequences are not submitted primary sequences. RefSeq records are owned by NCBI and therefore can be updated as needed to maintain current annotation or to incorporate additional sequence information." See also http://www.ncbi.nlm.nih.gov/LocusLink/refseq.html

It will be appreciated by those in the art that the exact results of a database search will change from day to day, as new sequences are added. Also, if you query with a longer version of the original sequence, the results will change. The results given here were obtained at one time and no guarantee is made that the exact same hits would be obtained in a search on the filing date. However, if an alignment between a particular query sequence and a particular database sequence is discussed, that alignment should not change (if the parameters and sequences remain unchanged).

Northern Analysis.

LO

<u>L</u>5

3 0

35

10

; 5

Northern analysis may be used to confirm the results. Favorable and unfavorable genes, identified as described above, or fragments thereof, will be used as probes in Northern hybridization analyses to confirm their differential expression. Total RNA isolated from subject mice will be resolved by agarose gel electrophoresis through a 1% agarose, 1 % formaldehyde denaturing gel, transferred to positively charged nylon membrane, and hybridized to a probe labeled with [32P] dCTP that was generated from the aforementioned gene or fragment using the Random Primed DNA Labeling Kit (Roche, Palo Alto, CA), or to a probe labeled with digoxygenin according to the manufacturer's instructions (Roche, Palo Alto, CA).

129

Real-Time RNA Analysis.

Real-time RNA analysis may also be used for confirmation. For "real-time" RNA analysis, RNA will be converted to cDNA and then probed with gene-specific primers made for each clone. "Real-time" incorporation of fluorescent dye will be measured to determine the amount of specific transcript present in each sample. Sample differences (older vs. younger) of 2-fold or greater (in either direction) will be considered differentially expressed. Confirmation using several independent animals is desirable.

In situ Hybridization

5

Another form of confirmation may be provided by nonisotopic in situ hybridizations (NISH) on selected human (obtained by Tissue Informatics) and mouse tissues using cRNA probes generated from mouse genes found to be up- or down-regulated during aging. In situ hybridizations may also be performed on mouse tissues using cRNA probes generated from differentially expressed DNAs. These cRNA's will hybridize to their corresponding messenger RNA's present in cells and will provide information regarding the particular cell types within a tissue that is expressing the particular gene as well as the relative level of gene expression. The cRNA probes may be generated by in vitro transcription of template cDNA by Sp6 or T7 RNA polymerase in the presence of digoxigenin-11-UTP (Roche Molecular Biochemicals, Mannheim, Germany; Pardue, M.L. 1985. situ hybridization, Nucleic acid hybridization, a practical approach: IRL Press, Oxford, 179-202).

Transgenic Animals.

Transgenic expression may be used to confirm the results. In one embodiment, a mouse is engineered to overexpress the favorable or unfavorable mouse gene in question. In another embodiment, a mouse is engineered to express the

130

corresponding favorable or unfavorable human gene. In a third embodiment, a nonhuman animal other than a mouse, such as a rat, rabbit, goat, sheep or pig, is engineered to express the favorable or unfavorable mouse or human gene.

5

10

15

3.0

25

30

:5

Hyperquantitative Tissue Analysis

In addition to gene expression analysis the tissue sections can also be analyzed using TissueInformatics, Inc's TissueAnalytics™ software. A single representative section may be cut from each tissue block, placed on a slide, and stained with H&E. Digital images of each slide may be acquired using an research microscope and digital camera (Olympus E600 microscope and Sony DKC-ST5). These images were acquired at 20x magnification with a resolution of 0.64 mm/pixel. A hyperquantitative analysis may be performed on the resulting images: First a digital image analysis can identify and annotate structural objects in a tissue using machine vision. These objects, that are constituents of the tissue, can be annotated because they are visually identifiable and have a biological meaning. (By way of example, for liver, the constituents can be, e.g., hepatocytes, sinusoids, vacuoles.) Subsequently a quantification of these structures regarding their geometric properties like area or stain intensities and their relationship to the field of view or per unit area in terms of a % coverage may be performed. Features or parameters for hyper-quantification are specific for each tissue, and may also include relations between features, measures of overall heterogeneity, including orientation, relative locations, and textures.

Correlation Analysis

Mathematical statistics provides a rich set of additional tools to analyze time resolved data sets of hyperquantitative and gene expression profiles for similarities, including rank correlation, the calculation of regression

131

and correlations coefficients, and clustering. Continuous functions may also be fitted through the data points of individual gene and tissue feature data. Relation between gene expression and hyper-quantitative tissue data may be linear or non-linear, in synchronous or asynchronous arrangements.

Introduction to Master Tables

)

The master tables reflect applicants' analysis of the gene chip data.

For each probe corresponding to a differentially expressed mouse gene, Master Table 1 identifies

- Col. 1: The mouse gene (upper) and mouse protein (lower) database accession #s.
- Col. 2: The corresponding mouse Unigene Cluster, as of the 4^{th} Quarter 2001 build.
- Col. 3: The behavior (differential expression) observed for the mouse gene. This column identifies the gene as favorable(F) or unfavorable (U) on the basis of its differential behavior in the comparisons (older vs. younger). As more than one older vs. younger comparison is made, only the result of the comparison yielding the greatest differential is listed. In the case of a gene with mixed behavior, both the result of the comparison yielding the greatest favorable differential and the result of the comparison yielding the greatest unfavorable differential are listed. If the value is followed by a parenthetical of the form "(X to Y)", it means that the differential value is the ratio when the absolute value for X weeks was compared to the absolute value for Y weeks, with the ratio being taken as greater-to-lesser.

132

One possible way of characterizing the degree of differential expression for a particular comparison would be to take the ratio of older to younger. If that ratio is at least 2:1, the behavior is considered unfavorable, and if it is not more than 0.5:1, it is unfavorable.

5

0

Use of an older/younger ratio is awkward when one wants to compare the degree of differential expression without regard to the direction of change. Consequently, in the Master Table, the numerical value is the ratio of the greater value to the lesser value. If this ratio is at least two fold, the degree of differential expression is considered significant.

In some of the related applications cited above, and perhaps occasionally in this application, a ratio may be given as a negative number. This does not have its usual mathematical meaning; it is merely a flag that in the comparison, the older value was less than the younger one, i.e., the gene was favorable. For the purpose of applying the teachings of the specification concerning desired ratios, any negative value should be converted to a positive one by taking its absolute value.

- Col. 4: A related human protein, identified by its database accession number. Usually, several such proteins are identified relative to each mouse gene. These proteins have been identified by BLAST searches, as explained in cols. 6-8.
- Col. 5: The name of the related human protein.
- Col. 6: The score (in bits) for the alignment performed by the BLAST program.
- Col. 7: The E-value for the alignment performed by the BLAST

133

program. It is worth noting that Unigene considers a Blastx E Value of less than 1e-6 to be a "match" to the reference sequence of a cluster.

5 Unless otherwise indicated, the bit score and E-value for the alignment is with respect to the alignment of the mouse DNA of col. 1 to the human protein of col. 4 by BlastX, according to the default parameters.

0

5

Master Table 1 is divided into two or three subtables on the basis of the Behavior" in col. 3. If a gene has at least one favorable behavior, and no unfavorable ones, it is put into Subtable 1A. In the opposite case, it is put into Subtable 1B. If any of the genes has mixed behavior, then Master Table 1 will include Subtable 1C for such genes.

Master Table 2 has just three columns.

- Col. 1: Mouse gene.
 - Col. 2: behavior. Same as col. 3 in Master table 1.
- Col. 3: Human protein classes. Based on the related human proteins defined in Master Table 1, Master Table 2 generalizes, if possible as to classes of human proteins which are expected to have similar behavior. For a given mouse gene, several human protein classes may be listed because of the diversity of the human proteins found to be related. In some cases, the stated human protein classes may be hierarchial, e.g., one may be a subset of another. In other cases, the stated classes may be non-overlapping but related. And in yet other cases, the stated classes may be non-overlapping and unrelated. Combinations of the above are also possible.

134

In addition to the classes stated, the corresponding human gene clusters are also of interest. These may be obtained in a number of ways. First, one may search on Unigene (http://www.ncbi.nlm.nih.gov/entrez/query.fcgi?db=unigene) for the identified human protein. Review the "hits" (each of which is a Unigene record) for those prefixed by "Hs." Secondly, one may access the Unigene record for the mouse gene cluster (which is given in Master Table 1), and then click on "Homologene". This will bring up a new page which includes the section "Possible Homologous Genes". One of the entries should be a Homo sapiens gene (considered by Unigene to be the most related human gene); click on its Unigene record link.

5

5

Additional information of interest may be accessed by searching with the mouse gene accession # in the Mouse Gene Informatics database, at http://www.informatics.jax.org/.

The related applications may contain reference to "2-16 week old mice". In the anti-diabetes series of applications, 3 week mice were put on a diet to induce obesity, hyperinsulinemia and diabetes. The 2-16 week old mice were more accurately described as mice who had been on that diet for 2-16 weeks, i.e., they were actually 5-19 weeks (35-133 days) old. Even some of the anti-aging series of applications made reference to 2-16 week old mice, even though the mice were in fact 5-19 weeks (35-133 days) old.

)) H		
Table 1A: Fa	ivorable G	Table 1A: Favorable Genes/Proteins				_
ŀ						
Mouse Gene	Gene Unigene	Behavior	Human	u m a n Description	Bits	E
Protein			Proteins			•
NM_008341 Mm.21300	Mm.21300	F:13.28 (5to11)	AAH35263.1	Similar to insulin-like growth factor binding protein 1	384	1E-106
NP 032367.1						
			CAA68770.1	CAA68770.1 IGF-binding preprotein (AA -25 to 234)	384	1E-106
			NP_000587.1	NP_000587.1 insulin-like growth factor binding protein 1	382	1E-106
			AAA52540.1	AAA52540.1 insulin-like growth factor binding protein precursor	338	9E-93
			CAA33110.1	CAA33110.1 small IGF-binding-protein	192	SE-50
NM_009669 Mm.324	Mm.324	F:8.34 (5to7)	NP_000690.1	000690.1 amylase, alpha 2A; pancreatic; Amylase, pancreatic, alpha-2A	952	0
NF 053799.1						
			NP_066188.1	_066188.1 amylase, alpha 2B; pancreatic; Amylase, pancreatic, alpha-2B	946	0
			XP_086988.1	086988.1 similar to Alpha-amylase, salivary precursor (1,4-alpha-D-glucan glucanohydrolase)	941	0
			67366	alpha-amylase (EC 3.2.1.1) precursor, salivary	939	0
			NP_004029.1	NP_004029.1 amylase, alpha 1A; salivary; Amylase, salivary, alpha-1A	939	0
			7245760	Chain A, Structure Of Human Pancreatic Alpha-Annylase In Complex With The Carbohydrate	927	0
				Inhibitor Acarbose		
			1421331	Chain, Mol_id: 1; Molecule: Human Pancreatic Alpha-Amylase; Chain: Null; Ec: 3.2.1.1	925	0
			18655894	Chain A, Three Dimensional Structure Analysis Of The R195q Variant Of Human Pancreatic	924	0
		-		Alpha Amylase		
			18655893	Chain A, Three Dimensional Structure Analysis Of The R337q Variant Of Human Pancreatic	924	0
				Alpha-Mylase	`,	-
		-	14719496	Chain A, Subsite Mapping Of The Active Site Of Human Pancreatic Alpha-Amylase Using	923	0
				Substrates, The Pharmacological Inhibitor Acarbose, And An Active Site Variant		

				1.36			
			20664071	Chain A, Mechanistic Analyses Of Catalysis In Human Pancreatic Alpha- Amylase: Detailed	923	0	
				Kinetic And Structural Studies Of Mutants Of Three Conserved Carboxylic Acids			
			20664068	Chain A, Three Dimensional Structure Analysis Of The R195a Variant Of Human Pancreatic	923	0	
				Alpha Anylase			
			18655892	Chain A, Three Dimensional Structure Analysis Of The R337a Variant Of Human Pancreatic	923	0	
				Alpha-Amylase			
			20664074	Chain A, Mechanistic Analyses Of Catalysis In Human Pancreatic Alpha- Amylase: Detailed	922	0	
·				Kinetic And Structural Studies Of Mutants Of Three Conserved Carboxylic Acids			
			1633119	Chain , Hunan Salivary Amylase	919	C	-,
			15988375	Chain A, Role Of Mobile Loop In The Mechanism Of Human Salivary Amylase	914	°	,
			15988376	Chain A, Role Of Ethe Mobile Loop In The Mehanism Of Human Salivary Amylase	904	0	_
		,	AAA57345.1	alpha-anylase	315	1E-146	
					910		
							-
NM_019824 Mm.24498		F:5.75 (7to19)	NP_005710.1	NP_005710.1 actin related protein 2/3 complex subunit 3; ARP2/3 protein complex subunit p21	365	1E-101	
NP 062798.1			*,				
			AAB61466.1 p21-Arc	p21-Arc	363	1E-100	
			*~				
			CAC14083.1	CAC14083.1 dJ470LJ4.3 (novel protein similar to the Arp2/3 protein complex subunit p21-Arc (ARC21))	350	8E-97	
			*~				
			XP_167194.1	similar to ARP2/3 complex 21 kDa subunit (P21-ARC) (Actin-related protein 2/3 complex	215	4E-56	
	-			subunit 3)			
NM_015763 Mm.28548		F:4.93 (5to19)	Q14693	Lipin 1	1493	0	
NP 056578.1							
			NP_663731.1 lipin 1		1488	-	_
			·			i	
			AAH30537.1	Similar to lipin 1	1487	0	
			XP 0411364		717		_
			, , , , , , , , , , , , , , , , , , ,	A CALLOS CONTROLL OF LEGICIES IN LANGE AND LOGIC CONTROLL OF LANGE AND LOGIC CONTROLL	14/6	0	

				137		
		·				
			NP_055461.1 lipin 2	lipin 2	790	0
NM_009117 Mm.148800 F:4.72 (5to19)	48800	F:4.72 (5to19)	NP_000322.1	NP_000322.1 serum amyloid A1	169	5E-43
NP 033143.1						
NM_015805 Mm.104687 F:4.48 (5to7)	04687	F:4.48 (5to7)	012110	Potential phospholipid-transporting ATPase IIA	1539	0
INF 050020.1						
			XP_030577.3	similar to Potential phospholipid-transporting ATPase IIA	1327	0
	-		BAA31586.1	BAA31586.1 KJAA0611 protein	1178	0
			CAB63450.1	CAB63450.1 [dJ1114A1.1 (ATPase, class II, type 9A (KIAA0611))	1133	0
			XP_085762.3	XP_085762.3 similar to ATPase, class 2, member b; ATPase 9B, class II; ATPase 9B, p type	658	P
			043861	Potential phospholipid-transporting ATPase IIB (HUSSY-20)	615	1E-175
·			AAC05243.1	putative ATPase	610	1E-174
			CAA06934.1	ATPase	609	1E-174
NM_007706 Mm.4132	132	F:4.4 (YtoM)	NP_003868.1	NP_003868.1 suppressor of cytokine signaling-2; STAT induced STAT inhibitor-2; cytokine-inducible SH2	364	1E-100
NP 031732.1				protein 2		
			JC5626	STAT induced STAT inhibitor 2	361	1E-100
			JC5760	cytokine-inducible SH2 protein 2	360	3E-99
			BAA22536.1	CIS2	359	3E-99
			AAC98896.1	suppressor of cytokine signalling-2; HSSOCS-2	350	3E-96
NM_008640 Mm.30071		F:4.09 (5to 19)	NP_055528.1	NP_055528.1 lysosomal-associated protein transmembrane 4 alpha; membrane nucleoside transporter;	390	1E-108
NF 032666.1				lysosomal-associated protein transmembrane 4		

1E-107	0	1E-180	4E-77	0		0	0	0		0	0	0	1E-131	1E-103	6E-70	2E-69
389	641	635	291	833		808	645	645		644	644	642	471	376	266	237
AAH03158.1 lysosomal-associated protein transmembrane 4 alpha	061821.1 Gene 33/Mig-6	Mig-6=mitogen-inducible gene mig-6 product [human, WI-38 cells, Peptide, 462 aa]	hypothetical protein DKFZp43411114.1	000679.1 aminolevulinate, delta-, synthase 1		5-arninolevulinate synthase precursor	5-aminolevulinic acid synthase	5-aminolevulinic acid synthase, erythroid-specific, milochondrial precursor (Delta-	aminolevulinate synthase) (Delta-ALA synthetase) (ALAS-E)	delta-aminolevulinate synthase (erythroid)	000023.1 aminolevulinate, delta-, synthase 2; Aminolevulinate, delta-, synthase-2	Similar to aminolevulinate, delta-, synthase 2 (sideroblastic/hypochromic anemia)	_115680.1 hypothetical protein MGC2605	similar to HAGH	_005317.1 hydroxyacyl glutathione hydrolase; hydroxyacyl glutathione hydroxyacyl glutathione hydroxyases glyoxalase II; hydroxyacylglutathione hydroxylase	
AAH03158.1	NP_061821.1	AAB35056.1	T46346	NP_000679.1		CAA68506.1	CAB06076.1	P22557		CAA39795.1	NP_000023.1	AAH30230.1	NP_115680.1	AAK61250.1	NP_005317.1	BAB70814.1
	F:4.06 (5to19)			F:3.98 (5to19)									F:3.89 (5to7)			,
				5 Mm.19143									Mm.195961			
	A K 0 0 4 8 5 1 Mm.21679	NF 396314.1		M 6 3 2 4 5	AAA91867.1				·				A K 0 0 5 2 7 4 Mm.195961 BAB23924.1			

10

ហ

			139		
NM_026346 Mm.40466 F:3.64 (YtoO)	F:3.64 (YtoO)	NP_478136	F-box only protein 32 isoform 1; muscle atrophy F-box protein; atrogin-1	710	0
NP 080622.1					
		BAB85128	F-box domain Fbx25-containing protein	446	1E-123
		NP 680482	F-box only protein 32 isoform 2; muscle atrophy F-box protein; atrogin-1	422	1E-116
		AAH24030	similar to RIKEN cDNA 4833442G10 gene (H. sapiens)	417	1E-115
		AAF04526	F-box protein Fbx25	354	5E-96
		NP 036305	F-box only protein 25; F-box protein Fbx25	353	6E-96
				·	
NM_025298 Mm.30605	F:3.45 (YtoM)	NP_060589	RNA polymerase III 80 kDa subunit RPC5	1288	0
NP 079574.1	-				
		AAM18215	RNA polymerase III 80 kDa subunit RPC5	1286	0
		BAB14481	lumamed protein product	1218	0
		AAH00285	hypothetical protein FLJ10509	1191	0
		BAB14437	unnamed protein product	1187	0
		BAA95976	KIAA1452 protein	1121	0
	•				
NM_022331 Mm.29151	F:3.44 (5to19)	NP_055500.1	NP_055500.1 homocysteine-inducible, endoplasmic reticulum stress-inducible, ubiquitin-like domain member	592	1E-168
NP 071726.1			1; MMS-inducible gene		
		AAC09357.1	имомп	525	1E-147
		AAG17233.1	unknown	295	2E-78
		AAH09739.1	Similar to homocysteine-inducible, endoplasmic reticulum stress-inducible, ubiquitin-like domain	218	2E-55
			member 1		
		NP_071768.2	NP_071768.2 hypothetical protein FLJ22313	216	1E-54
NM_016773 Mm.9901	F:3.41 (YtoO)	NP_005004.1	NP_005004.1 nucleobindin 2	675	0
NP 058053.1					
		AAM73810.1	AAM/73810.1 NUCB2 protein	699	0

٠					
		NP_006175.2	NP_006175.2 nucleobindin 1	424	1E-118
		Q02818	Nucleobindin 1 precursor (CALNUC)	418	1E-116
		AAA36383.1 nucleobindin	nucleobindin	414	1E-115
B C 0 1 7 6 0 3 Min.204670 F:3.36 (5to19)	7:3.36 (51019)	BAC11593.1	unnamed protein product	384	1E-105
AAH17603.1					
		NP_110382.1	NP_110382.1 thioredoxin-related transmembrane protein	383	1E-105
		T12471	hypothetical protein DKFZp564E1962.1	383	1E-105
	-	AAH36460.1	Similar to thioredoxin domain-containing	381	1E-104
			hypothetical protein DJ971N18,2	202	1E-50
		BAC11237.1	unnamed protein product	202	1E-50
	-	CAC17521.1	dJ971N18.2.1 (novel protein (isoform 1))	202	1E-50
		·			
NM_013584 Mm.3174 F NP_038612.1	F:3.35 (5to 19)	NP_002301.1	NP_002301.1 leukemia inhibitory factor receptor precursor	1663	0
		AAB23884.1	AAB23884.1 leukaemia inhibitory factor receptor, LIF receptor [human, placenta, Peptide, 1078 aa]	1640	0
		NP_003990.1	oncostatin M receptor	345	2E-94
		AAB61897.1	leukemia inhibitory factor receptor	282	2E-75
NM_013590 Mm.177539 F:3,34 (7t019) NP_038618.1	F:3,34 (7to19)	NP_000230,1	NP_000230.1 lysozyme precursor	233	6E-62
		AAA36188.1	lysozyme precursor (EC 3.2.1.17)	233	2E-61
		AAC63078.1	lysozyıne precursor	228	2E-60
				-	

S

10

 $\vec{\circ}$

			1.41		. [
		pdb 1C7P	Chain A, Crystal Structure Of Mutant Human Lysozyme With Four Extra Residues (Eaea) At The	228	2E-60
			N-Terminal		
		pdb 1B7P	Chain A, Verification Of Sprap Using Mutant Human Lysozymes	228	3E-60
A K 0 0 5 5 4 6 Mm.33326 F.	F:3.33 (5to19)	NP_000119.1	_000119.1 plasma coagulation factor XI precursor, isoform a; plasma thromboplastin antecedent	1048	0
		1 1 0 0 0 0 0 1	124	104	(
		AAA51985.1	AAA51985.1 coagulation factor XI	1044	0
		NP_062505.1	062505.1 platelet coagulation factor XI, isoform b; plasma thromboplastin antecedent	848	0
		NP_000883.1	NP_000883.1 plasma kallikrein B1 precursor; Kallikrein, plasma; kallikrein 3, plasma; kallikrein B plasma;	755	0 ·
			Fletcher factor		
		AAC51784.1	serine protease	217	5E-55
		AAK53559.1	epitheliasin	216	1E-54
		AAK29280.1	androgen-regulated serine protease TMPRSS2 precursor	216	1E-54
		NP_005647.2	_005647.2 transmembrane protease, serine 2; epitheliasin	216	1E-54
NM_010286 Mm.22216 F	F:3.32 (5to19)	99576	Glucocorticoid-induced leucine zipper protein (Delta sleep-inducing peptide innunoreactor)	196	8E-49
_			(DSP-immunoreactive peptide) (DIP protein) (hDIP) (TSC-22-like protein) (TSC-22R)		
		T14749	hypothetical protein DKF2p566A093.1	188	2E-46
		-			
NM_009344 Mm.3117 F	F:3.29 (7to19)	NM_009344	NM_009344 pleckstrin homology-like domain, family A, member 1; PQ-rich protein	379	1E-104
	-	NP_033370.1			
		AAH18929.1	Similar to T-cell death associated gene	235	1E-60

WO 2005/000335

142

ហ

0		1E-172	0		0		0	0	0	0	0	0		1E-165	1E-165	1E-165	1E-164	1E-163	1E-158	1E-153	1E-152
700		578	1253		1249		1238	1179	1151	981	749	746		185	581	185	579	574	556	540	539
NP_570901.1 solute carrier family 39 (zinc transporter), member 4		solute carrier family 39 (zinc transporter), member 4	E-cadherin		004351.1 cadherin 1, type 1 preproprotein; calcium-dependent adhesion protein, epithelial; cadherin 1, E-	cadherin (epithelial); uvomorulin; cell-CAM 120/80; Arc-1		E-cadherin	AAA612591 uvomorulin	E-cadherin	Cadherin-3 precursor (Placental-cadherin) (P-cadherin)	001784.2 cadherin 3, type 1 preproprotein; P-cadherin; placental cadherin; cadherin 3, P-cadherin	(placental); calcium-dependent adhesion protein, placental	Neural-cadherin precursor (N-cadherin) (Cadherin-2)	NP_001783.2 cadherin 2, type 1 preproprotein; N-cadherin 1; cadherin 2, N-cadherin (neuronal); neural	N-cadherin	cadherin 2 precursor	H36470.1 cadherin 2, type 1, N-cadherin (neuronal)	_001785.2 cadherin 4, type 1 preproprotein, cadherin 4, R-cadherin (retinal); R-cadherin; retinal cadherin	Cadherin-4 precursor (Retinal-cadherin) (R-cadherin) (R-CAD)	N-cadherin
NP_570901.		NP_060237.1	CAA79356,1		NP_004351.		BAA88957.1	CAA84586.1	AAA61259	BAA88956.1	P22223	NP_001784.		P19022	NP_001783.	AAB22854.1	IJHUCN	AAH36470.1	NP_001785	P55283	AAA03236.1
F:3.25 (YtoM)			F:3.24 (YtoO)				,									- -					
Mm.29483																					
A K 0 0 5 5 3 5 Mm 29483	BAB24106.1		NM_009864 Mm.35605	NP 033994.1			•														

		CAA40773.1	N-cadherin	526	1E-148
		BAC03677.1	unnamed protein product	523	1E-147
	1				
NM_007687 Mm.4024 F NP_031713.1	F:3.24 (5to7)	NP_005498.1		327	1E-88
		NP_068733.1	NP_068733.1 cofilin 2 isoform 1	274	IE-72
		XP_171270.1	XP_171270.1 similar to COFILIN, NON-MUSCLE ISOFORM	261	1E-68
		XP_060203.1	_060203.1 similar to Cofilin, non-muscle isoform (18 kDa phosphoprotein)(P18)	248	8E-65
		NP_006861.1	006861.1 destrin (actin depolymerizing factor); destrin	237	3E-61
A K 0 0 5 9 8 9 Mm.182959 F:3.2 (YtoM) BAB24354.1	F:3.2 (YtoM)	NP_005733.1		714	0
		AAB50217.1	B50217.1 protein disulfide isomerase-related protein 5	681	0
,					T
NM_008904 Mm.10707 F	F:3.2 (YtoM)	AAD51615	PPAR gamma coactivator-1	1273	0
		NP 037393	peroxisome proliferative activated receptor, gamma, coactivator 1	1272	0
NM_026508 Mm.182051 F:3.15 (YtoM) NP_080784.1	F:3.15 (YtoM)	NP_057376.1	_057376.1 tumor necrosis factor type 1 receptor associated protein	1214	0
	·	AAH18950.1	Unknown (protein for MGC:15157)	1214	0
		AAH01455.1	H01455.1 heat shock protein 75	1208	0
		AAC02679.1	C02679.1 heat shock protein 75	1144	0

		A55877	tumor necrosis factor type 1 recentor associated protein TRAP-1	1108	-
		AAC24722.1	TRAP1	1082	0
		BAC04139.1	BAC04139.1 unnamed protein product	066	0
		AAH02994.1	AAH02994.1 Unknown (protein for MGC;3823)	741	0
		AAK74072.1	AAK74072.1 heat shock protein gp96 precursor	324	3E-88
		NP_003290.1	NP_003290.1 tumor rejection antigen (gp96) 1; Tumor rejection antigen-1 (gp96)	323	7E-88
NM_021792 Mm.29008 F:3.14 NP 068564.1	F:3.14 (5to19)	NP_062558.1	NP_062558.1 hypothetical protein R30953_1	229	2E-59
N.M025404 Mm.5376 F:3.11 NP_079680.1	F:3.11 (50011)	AAH00043.1	ADP-ribosylation factor 4-like	359	6E-98
		NP_001652.1	NP_001652.1 ADP-ribosylation factor 4-like; ADP-ribosylation factor-like 6	357	2E-97
		AAA93229.1	ADP-ribosylation factor	348	1E-94
		XP_045890.2	XP_045890.2 similar to ADP-ribosylation factor 4L.	233	4E-60
		NP_005729.1	NP_005729.1 ADP-ribosylation factor-like 4	209	9E-57
		NP_005728.2	NP_005728.2 ADP-ribosylation factor-like 7	148	1E-52
A K 0 0 5 0 3 5 Mm.18802 F:3.09 BAB23762.1	F:3.09 (5to 19)	NP_001054.1 transferrin	transferrin	853	0
		pdb 1LFG	Lactoferrin (Diferric)	797	0
		pdb 1LFH	Lactoferrin (Apo Form)	797	0
		pdb 1LFI	Lactofernin (Copper Form)	797	0
		pdb 1FCK	Lactoferrin	<i>16L</i>	0

WO 2005/000335 PCT/US2004/017322

		145		
	AAA59479.1	AAA59479.1 neutrophil lactoferrin	962	0
	pdb 1B0L	Lactoferrin	961	0
	pdb/1CB6	Lactoferrin	962	0
	CAA37116.1	precursor lactoferrin (709 AA)	962	0,
	P02788	Lactotransferrin precursor (Lactoferrin) [Contains: Lactoferroxin A; Lactoferroxin B;	962	0
		Lactoferroxin C]		
	pdb 1LCF	pdb 1LCF Lactoferrin (Copper and Oxalate Form)	96L	0
·	AAH15822.1	lactotransferrin	796	0
	AAA36159.1	lactoferrin	795	0
	AAA58656.1	III.F2	795	0
	AAH22347.1	lactotransferrin	795	0
	AAG48753.1	AAG48753.1 lactoferrin precursor	794	0
	AAA59511.1	lactoferrin	793	0
-	AAH15823.1	AAH15823.1 lactotransferrin	793	0
	NP_002334.1	002334.1 lactotransferrin	792	0
	AAF22007.1	PRO1400	709	0
	AAA61141.1	transferrin	622	1E-177
	AAB57795.1	AAB57795.1 Lactoferrin. Incomplete at NH2 end	570	570 1E-161

10 15

ហ

			146		
		XP_067170.1	067170.1 similar to RIKEN cDNA 1300017J02	550	1E-155
		pdb 1FQF	Serotransferrin	443	1E-123
		pdb/1BP5	Serum Transferrin	443	1E-123
		pdb 1FQE	Serotransferrin	441	1E-122
		pdb[1A8E	Serum Transferrin	440	1E-122
		pdb/1.1QF	Transferrin	440	1E-122
		pdb 1D3K	Serum Transferrin	439	1E-121
		pdb 1DTG	Transferrin	438	1E-121
		pdb 1B3E	Serum Transferrin	437	IE-121
		pdb 1D4N	Transferrin	437	1E-121
		NP_005920.1	005920.1 melanoma-associated antigenp97, isoform 1, precursor; melanotransferrin; melanoma-associated	394	1E-108
			autigen p97		
		pdb 1LCT	Lactoferrin (N-Terminal Half-Molecule)	372	1E-101
		pdb 1EH3	Lactoferrin	372	1E-101
		pdb 1L5T	Lactoferrin	370	1E-101
		pdb 1VFE	Human Lactoferrin	370	1E-101
		pdb 1DSN	Lactoferrin	368	1E-100
		pdb 1VFD	Lactoferrin	367	1E-100
83 Mm.4863	F:3.09 (5to19)	CAC14276.1	CAC14276.1 bA112L6.1 (CCAAT/enhancer binding protein (C/EBP), beta)	271	2E-72
NP 034013.1					
		NP_005185.1	005185.1 CCAAT/enhancer binding protein (C/EBP), beta; CCAAT/enhancer-binding protein (C/EBP),	271	2E-72
			beta (transcription factor-5)		
01 Min.63479	F:3.08 (YtoM)	AAH44572	similar to solute carrier family 15 (H+/peptide transporter), member 2	1128	0
NP 067276.1					
		NP 066568	solute carrier family 15 (H+/peptide transporter), member 2	1122	
		AAC15477	Caco-2 oligopeptide transporter	561	1E-159

	:		147		
		NP_005064	solute carrier family 15 (oligopeptide transporter), member 1; Human peptide transporter	195	1E-159
			(HPEPT1) mRNA, complete cds	·	
		CAC27442	bA551M18.1.1 (solute carrier family 15 (oligopeptide transporter) member 1)	502	1E-141
		JC5638.	pH-sensing regulatory factor	231	5E-60
NM_013786 Mm.26719	9 F:3.08 (YtoM)	NP_003716	3-hydroxysteroid epimerase; oxidative 3-alpha-hydroxysteroid-dehydrogenase; 3(alpha->beta)-	473	1E-133
NP_038814.1			hydroxysteroid epimerase; retinol dehydrogenase; oxidoreductase; NAD+ -dependent 3 alpha-		•
			hydroxysteroid dehydrogenase		
		AAB88252	oxidative 3 alpha hydroxysteroid dehydrogenase	442	1E-124
		AAC39922	sterol/retinol dehydrogenase	404	1E-112
		NP 003699	microsomal NAD+-dependent retinol dehydrogenase 4	399	1E-111
		NP 683695	orphan short-chain dehydrogenase / reductase; retinol dehydrogenase similar prolein	312	7E-85
		NP 005762	NADP-dependent retinol dehydrogenase/reductase; 3-alpha hydroxysteroid dehydrogenase	300	3E-81
		Q92781	11-cis retinol dehydrogenase (11-cis RDH).	283	4E-76
,		AAH28293	Similar to retinol dehydrogenase 5 (11-cis and 9-cis)	281	1E-75
		NP 002896	retinol dehydrogenase 5 (11-cisand 9-cis); retinol dehydrogenase 5 (11-cis and 9-cis)	272	1E-72
		AAD32458	retinol dehydrogenase homolog	267	2E-71
		AAF82748	retinol dehydrogenase homolog isoform-1	252	1E-66
NM_016917 Mm.28756	6 F:3.08 (7to19)	NP_055400.1	NP_055400.1 solute carrier family 11 (proton-coupled divalent metal ion transporters), member 3; ferroportin	936	0
NP 058613.1			1; iron regulated gene 1; ferroportin 1		
		AAF80986.1	SLC11A3 iron transporter	933	0
		AAH35893.1	solute carrier family 11 (proton-coupled divalent metal ion transporters), member 3	932	0
-		BAA92049.1		302	2E-81
NM_010004 Mm.29973	73 F:3.08 (5to19)	NP_000763.1	NP_000763.1 cytochrome P450, subfamily IIC (mephenytoin 4-hydroxylase), polypeptide 18; cytochrome	629	1E-179
NP_034134.1			P450, subfamily IIC (mephenytoin 4-hydroxylase), polypeptide 17; microsomal monooxygenase;		
			flavoprotein-linked monooxygenase		
		P33260	Cytochrome P450 2C18 (CYPIIC18) (P450-6B/29C)	627	1E-178

Ŋ

	NP 000760.1	000760.1 cytochrome P450, subfamily IIC (mephenytoin 4-hydroxylase), polypeptide 19; mephenytoin 4'-	618	618 1E-176
	l	hydroxylase; microsomal monooxygenase; xenobiotic monooxygenase; flavoprotein-linked		•
		топоохуденаѕе		
	F38462	S-niephenytoin 4'-hydroxylase (EC 1.14.14) cytochrome P450 2C19	612	1E-174
	AAB23864.2	.B23864.2 cytochrome P-450	611	1E-174
	BAA00123.1	cytochrome P-450	611	1E-174
	1506290A	cytochrome P450	610	1E-173
	P11712	Cytochrome P450 2C9 (CYPIIC9) (P450 PB-1) (P450 MP-4) (S-mephenytoin 4-hydroxylase)	611	1E-174
		(P-450MP).		
	AAA52157.1	cytochrome P-450 S-mephenytoin 4-hydroxylase	605	1E-172
	P10632	Cytochrome P450 2C8 (CYPIIC8) (P450 form 1) (P450 MP-12/MP-20) (P450 IIC2) (S-	009	1E-170
		mephenytoin 4-hydroxylase)		
	AAH20596.1	AH20596.1 Unknown (protein for MGC:22146)	599	1E-170
	AAA52161.1	cytochrome P-450 S-mephenytoin 4-hydroxylase	598	1E-170
	AAB35292.1	AAB35292.1 cytochrome P450 arachidonic acid epoxygenase isoform, Cyp 2C8 [human, kidney, Peptide	969	1E-169
		Partial, 485 aa]		
	AAA36660.1	AAA36660.1 cytochrome P450	598	1E-170
	AAA35739	cytochrome P-450 1	597	1E-169
	AAB35292.1	cytochrome P450 arachidonic acid epoxygenase isoform, Cyp 2C8 [human, kidney, Peptide	965	1E-169
 		Partial, 485 aa].		
	AAA52160	cytochrome P-450 S-mephenytoin 4-hydroxylase.	595	1E-169
	AAA52145.1	locus HUMCYP2C17	516	1E-145
	AAA52159.1	cytochrome P-450 S-mephenytoin 4-hydroxylase	514	1E-144
	CAA46778.1	CAA46778.1 cytocluome P-450 II C	497	1E-139

			149		
		A31949	cytochrome P450 2B1	478	1E-133
		AAD13753.1	cytochrome P450 2E1	478	1E-133
		AAF13601.1	cytochrome P450-2E1	478	1E-133
A B 0 6 0 2 7 4 Mm. 196225 F:3.04 (YtoM) BAB41208.1	.25 F:3.04 (YtoM)	NP_001944	endothelial cell growth factor 1 (platelet-derived); thymidine phosphorylase; gliostatin	730	0
		AAH18160	endothelial cell growth factor 1 (platelet-derived)	728	C
		P19971	Thymidine phosphorylase precursor (TdRPase) (TP) (Platelet-derived endothelial cell growth	728	0
	·		factor) (PD-ECGF) (Gliostatin).		-
		NP 005129	cytochrome oxidase deficient homolog 2	234	3E-61
		NP 004580	cytochrome oxidase deficient homolog 1	234	3E-61
NM_018887 Mm.17991	11 F:3 (7to19)	NP_057677.1	057677.1 oxysterol 7alpha-hydroxylase	712	0
NP 061375.1					
		AAH10358.1	oxysterol 7alpha-hydroxylase	710	0
NM_024406 Mm.582	F:2.98 (7to19)	NP_001433.1	_001433.1 fatty acid binding protein 4, adipocyte; A-FABP	245	4E-65
NP 077717.1	-	·			
NM_018746 Mm.34819	19 F:2.96 (YtoM)	BAA07536.1	BAA07536.1 PK-120 precursor	1130	0
NP 061216.1	·				
		AAD05198.1	inter-alpha-trypsin inhibitor family heavy chain-related protein	1127	0
		Q14624	Inter-alpha-trypsin inhibitor heavy chain H4 precursor (ITI heavy chain H4) (Inter-alpha-inhibitor	1127	0
			heavy chain 4) (Inter-alpha-trypsin inhibitor family heavy chain-related protein) (IHRP) (Plasma		
			kallikrein sensitive glycoprotein 120) (PK-120) (GP120) (PRO1851) [Contains: GP57]		
		NP_002209.1	NP_002209.1 inter-alpha (globulin) inhibitor H4 (plasma Kallikrein-sensitive glycoprotein); inter-alpha	1126	0
			(globulin) inhibitor, H polypeptide-like 1; Inter-alpha (globulin) inhibitor, H4 polypeptide		
		AAF69610.1	PRO1851	722	0

ט

 \Box

8.1 I I I I I I I I I I I I I I I I I I I						
S30350 1						
NP_002208.1			350	inter-alpha-trypsin inhibitor heavy chain 3 precursor	0 79	1E-177
P19827 P19827 P19821 P19823 P19			002208.1	pre-alpha (globulin) inhibitor, H3 polypeptide; Inter-alpha (globulin) inhibitor, H3 polypeptide	620	1E-177
S24391 NP_002206.11 P19823 P19823 P19823 P19823 P19823 P19823 P19823 P19823 P19823 P1002207.1 P100207.1 P1002007.1 P2001697.2 P2001697.2 PA48752 PA48752 PA48752 PA48752 PA48752 PA48752 PA48752 PA48752 PA600962.1 P2000230.1 Pablic7P				Inter-alpha-trypsin inhibitor heavy chain H1 precursor (ITI heavy chain H1) (Inter-alpha-inhibitor	461	1E-129
S24391				$heavy\ chain\ 1) (Inter-alpha-tryps in inhibitor\ complex\ component\ III) (Serum-derived\ hyaluronan-derived\ hyaluronan-de$		
S24391 ii NP_002206.1 ii NP_002206.1 ii NP_002207.1 ii NP_002207.1 ii NP_002207.1 ii NP_001697.2 ii NP_01697.2 ii NP_01697.1 ii				associated protein) (SHAP)		
NP_002206.1 P19823 P19823 P19823 P202207.1 P20			391	inter-alpha-trypsin inhibitor heavy chain H1 precursor	461	1E-129
P19823 NP_002207.13 S04484 CAA34346.1 CAA34346.1 A48752 BAC00962.1 XP_171849.1 XP_171849.1 AAA36188.1 P002207.1 P002207.1 P001697.2 PAA36188.1			NP_002206.1	inter-alpha (globulin) inhibitor, H1 polypeptide	461	1E-129
NP_002207.1 S04484 S04484 CAA34346.1 H4 Mm.15811 F.2.93 (5to19) NP_001697.2 BAC00962.1 XP_171849.1 XP_171849.1 AAA36188.1 Pabbl1C7P				Inter-alpha-trypsin inhibitor heavy chain H2 precursor (ITI heavy chain H2) (Inter-alpha-inhibitor	449	1E-125
NP_002207.13 \$04484 \$04484 CAA34346.1 CAA34346.1 P_001697.2 BAC00962.1 XP_171849.1 XP_171849.1 XP_171849.1 AAA36188.1				$\it heavy chain 2) (Inter-alpha-tryps in inhibitor complex component II) (Serum-derived hyaluronan-derived hyaluronan-derived$		
NP_002207.1 S04484 S04872 S04484 S04872 S04484 S04872 S044872 S044872 S04484 S04872 S04484 S04872 S04484 S04872 S04872 S04972 S0				associated protein) (SHAP)		
14 Mm.15811 F:2.93 (5to19) NP_001697.2 A48752 BAC00962.1 XP_171849.1 XP_171849.1 AAA36188.1			NP_002207.1	inter-alpha (globulin) inhibitor, H2 polypeptide	447	1E-125
14 Mm.15811 F:2.93 (5to19) NP_001697.2 A48752 BAC00962.1 XP_171849.1 AAA36188.1				inter-alpha-trypsin inhibitor chain 3 - human	441	1E-123
14 Mm.15811 F:2.93 (5to19) NP_001697.2 A48752 BAC00962.1 XP_171849.1 72 Mm.45436 F:2.91 (7to19) NP_000230.1			CAA34346.1	inter-alpha-trypsin inhibitor C-terminal	434	1E-121
14 Mm.15811 F:2.93 (5to19) NP_001697.2 A48752 BAC00962.1 XP_171849.1 72 Mn.45436 F:2.91 (7to19) NP_000230.1						
A48752 BAC00962.1 XP_171849.1 72 Mm.45436 F:2.91 (7to19) NP_000230.1 AAA36188.1	14 Mm.15811	:2.93 (5to19)		B-cell lymphoma 6 protein; B-cell CLL/lymphoma-6; cys-his2 zinc finger transcription factor	1337	0
BAC00962.1 XP_171849.1 F:2.91 (7to19) NP_000230.1 AAA36188.1	14F 0536/4:1		A48752	B-cell CLI Avmohoma 6 (BCL6) protein	1330	70
XP_171849.1 F:2.91 (7to19) NP_000230.1 AAA36188.1			62.1	BAZF	335	1E-91
F:2.91 (7to19) NP_000230.1 AAA36188.1				similar to BcL6-associated zinc finger protein	300	7E-81
F:2.91 (7to19) NP_000230.1 AAA36188.1 pdbl1C7P						
AAA36188.1		:2.91 (7to19)		lysozyme precursor	231	3E-61
			AAA36188.1	lysozyme precursor (EC 3.2.1.17)	230	5E-61
٦			pdb 1C7P	Chain A, Crystal Structure Of Mutant Human Lysozyme With Four Extra Residues (Eaea) At The	228	3E-60

2.

3.0

3

. ザ

				151		
				N-Terminal		
			pdb 1B7P	Chain A, Verification Of Spmp Using Mutant Human Lysozymes	227	6E-60
			pdb 133L	Chain, Lysozyme (E.C.3.2.1.17) Mutant With Arg 115 Replaced By His (R115h)	227	6E-60
			AAC63078.1	lysozyme precursor	226	1E-59
NM_021313Mm.86910 NP 067288.1	Mm.86910	F:2.91 (5to19)	096ВН1	RING finger protein 25	702	0
-			NP_071898.1	NP_071898.1 ring finger protein 25	684	0
X93035 CAA63603.1	Mn.4376	F.2.87 (YtoO)	AAH08568	Similar to chitinase 3-like 1 (cartilage glycoprotein-39)	537	1E-152
			NP 001267	chitinase 3-like 1; cartilage glycoprotein-39	536	1E-152
			AAH38354	similar to chitinase 3-like 1 (cartilage glycoprotein-39)	535	1E-152
			NP 003456	chitotriosidase; plasma methylumbelliferyl tetra-N-acetylchitotetraoside hydrolase	355	8E-98
			AAG10644	chitotriosidase precursor	355	1E-97
			ILGIA.	Chain A, Crystal Structure Of Human Chitotriosidase In Complex With Chitobiose	345	8E-95
			NP 003991	chitinase 3-like 2; chondrocyte protein 39	340	3E-93
			Q15782	Chitinase 3-like protein 2 precursor (YKL-39) (Chondrocyte protein 39).	340	3E-93
			AAH11460	chitinase 3-like 2	340	3E-93
			AAB04534	chitinase	340	3E-93
			AAG60019	acidic mammalian chitinase precursor	319	8E-87
			AAO37816	oviductin	274	2E-73
			AAB04126	oviductal glycoprotein	273	5E-73
			NP_002548	oviductal glycoprotein 1, 120kDa (mucin 9, oviductin); mucin 9 (oviductin); oviductal	273	5E-73
				glycoprotein 1, 120kD (mucin 9, oviductin)		-
			138605	oviductal glycoprotein	273	5E-73
			NP 068569	eosinophil chemotactic cytokine	226	9E-59
			,			
NM_023184 Mm.41389	Mm.41389	F:2.87 (5to11)	NP 054798.1	NP_054798.1 Kruppel-like factor 15; KKLF protein; kidney-enriched Kruppel-like factor	624	1E-178
NP 075673.1						

52

Ŋ

	2E-56	2E-82	8E-82		3 2E-80	5 8E-74		4E-72	3 2E-65	0			0		0 /	1 1E-180	1E-134	3 1E-132	3 1E-132	3 1E-132
	220	305	302		298	276		270	248	1392			1392		637	634	479	473	473	473
752	NP_001435.1 fatty acid binding protein 5 (psoriasis-associated); E-FABP	BAC11635.1 unnamed protein product	Osteopontin precursor (Bone sialoprotein 1) (Urinary stone protein) (Secreted phosphoprotein	1) (SPP-1) (Nephropontin) (Uropontin)	OPN-a - human (fragment).	secreted phosphoprotein 1 (osteopontin, bone sialoprotein I, early T-lymphocyte activation 1);	Secreted phosphoprotein-1 (osteopontin, bone sialoprotein)	OPN-b - human (fragment).	OPN-c - human (fragment).	NP_005202.1 colony stimulating factor 1 receptor, formerly McDonough feline sarcoma viral (v-fms) oncogene	homolog; C-FMS, Colony-stimulating factor-1 receptor; oncogene FMS (McDonough feline	sarcoma)	Macrophage colony stimulating factor I receptor precursor (CSF-1-R) (Fms proto-oncogene) (c-	fms) (CD115 antigen)	NP_000213.1 v-kit Hardy-Zuckerman 4 feline sarcoma viral oncogene homolog precursor	KIT protein	NP_006197.1 platelet-derived growth factor receptor alpha precursor	NP_002600.1 platelet-derived growth factor receptor beta precursor; beta platelet-derived growth factor recentor		AAA36427.1 platelet-derived growth factor receptor.
	NP_001435.1	BAC11635.1	P10451		156986	NP_000573.1		176601	176602	NP_005202.			P07333		NP_000213.	AAC50969.1	NP_006197.	NP_002600.	AAH32224.1	AAA36427.1
	F.2.84 (5to19)	F.2.82 (5to19)								F:2.8 (5to 19)				·						
		Mm.321																		
	NM_010634 Mm.741 NP 034764.1	NM_009263 Mm.321 NP 033289.1								NM_007779 Mm.22574	NP_031805.1						`			

			153		
		CAA81393.1	CAA81393.1 FLT3 receptor tyrosine kinase	419	1E-115
		NP_004110.1	004110.1 fms-related tyrosine kinase 3	416	1E-114
		A36873	protein-tyrosine kinase (EC 2.7.1.112) STK-1 precursor	408	1E-112
NM_011825 Mm.25760 NP_035955.1	F:2.8 (5to19)	NP_071914.1	_071914.1 hypothetical protein FLJ21195 similar to protein related to DAC	308	5E-83
		NP_037504.1	037504.1 cysteine knot superfamily 1, BMP antagonist 1; gremlin	184	1E-45
A K 0 0 7 7 0 7 Mm.9806	F:2.79 (YtoO)	AAF64142	NPD008 protein	394	1E-109
BAB25202.1					
		AAH08430	Unknown (protein for MGC:14598)	391	1E-108
		NP 057162	CGI-148 protein	349	7E-96
		NP 660344	similar to CGI-148 protein	293	SE-79
NM_026007 Mm.42960) F:2.76(YtoM)	NP_001395.1	001395.1 eukaryotic translation elongation factor 1 gamma; elongation factor 1-gamma; EF-1-gamma; eEF-	792	0
NP_080283.1			1B gamma; translation elongation factor eEF-1 gamma chain; PRO1608; pancreatic tumor-related		
			protein		
		AAH13918.1	Similar to cukaryotic translation elongation factor 1 gamma	16/	0
		XP_088122.2	. 088122.2 similar to Elongation factor 1-gamma (EF-1-gamma) (eEF-1B gamma)	779	0
-		AAC18414.1	pancreatic tumor-related protein	99	0
		AAF69604.1 PRO1608	PRO1608	280	1E-165
NM_024169 Mm.30729	9 F:2.76 (5to19)	NP_057678.1	057678.1 FK 506 binding protein precursor; FK 506 binding protein 11 (19 kDa)	294	1E-79
NP 077131.2					
NM 008061 Mm. 18064	4 F:2.75 (5to11)	NP 000142.1	000142.1 glucose-6-phosphatase, catalytic	588	1E-168

LO

0

15

 $^{\circ}$

ζ, (Σ)

				154		
NP_032087.1						
			AAH20700.1	AAH20700.1 Unknown (protein for MGC:22459)	416	1E-115
			NP_066999.1	islet-specific glucose-6-phosphatase catalytic subunit-related p	318	2E-86
NM_019806 Mm.38800	Mm.38800	F:2.74 (5to19)	NP_004729.1	NP_004729.1 VAMP (vesicle-associated membrane protein)-associated protein B and C; VAMP-associated	404	1E-113
NP_062780.1				protein C; VAMP-associated protein B; VAMP-associated 33 kDa protein		
			AAF67013.1	VAMP-associated 33 kDa protein	399	1E-111
			AAF72105.1	33 kDa Vamp-associated protein	291	SE-79
			NP_003565.2	NP_003565.2 vessicle-associated membrane protein (VAMP)-associated protein of 33 kDa; vesicle-associated	291	8E-79
:				membrane protein (VAMP), 33 kDa; VAMP-associated protein A; VAMP (vesicle-associated		
				membrane protein)-associated protein A (33kD)		
			AAC26508.1	AAC26508.1 VAMP-associated protein of 33 kDa	289	3E-78
NM_022324 Mm.30222 NP_071719.1	Mm.30222	F:2.74 (5to19)	AAH10880.1	AAH10880.1 Unknown (protein for MGC:1757)	342	5E-94
			AAH06248.1	AAH06248.1 stromal cell-derived factor 2-like 1	340	3E-93
			NP_071327.1	NP_071327.1 stromal cell-derived factor 2-like 1	334	1E-91
			NP_008854.2	NP_008854.2 stromal cell-derived factor 2 precursor	236	3E-62
			Q99470	Stromal cell-derived factor 2 precursor (SDF-2)	233	5E-61
M 1 2 5 7 1	Mm.196559	2 5 7 1 Mm.196559 F:2.73 (YtoM)	NP_005336.2	NP_005336.2 heat shock 70kDa protein 1A; heat shock 70kD protein 1A; heat shock-induced protein; dnaK-	635	0
AAA57234.1				type molecular chaperone HSP70-1		
			P08107	Heat shock 70 kDa protein 1 (HSP70.1) (HSP70-1/HSP70-2)	635	0

5

 $\vec{\vdash}$

5

 $\frac{1}{2}$

WO 2005/000335 PCT/US2004/017322

155

		NP_005337.1	NP_005337.1 heat shock 70kDa protein 1B; heat shock 70kD protein 1B	633	1E-180
		A29160	A29160 dnaK-type molecular chaperone HSPA1L	628	1E-179
		XP_175177.1	heat shock 70kD protein 1-like	588	1E-167
		BAA32521.1	Heat shock protein 70 testis variant	989	1E-166
		NP_005518.1	NP_005518.1 heat shock 70kDa protein 1-like; Heat-shock 70kD protein-like-1; heat shock 70kD protein-like	L	586 IE-166
		XP_166348.1	XP_166348.1 similar to Heat shock 70 kDa protein 1-HOM (HSP70-HOM)	989	1E-166
		AAH34483.1	AAH34483.1 heat shock 70kD protein 1-like	585	IE-166
		NP_068814.2	NP_068814.2 heat shock 70kDa protein 2; heat shock 70kD protein 2; Heat-shock 70kD protein-2	567	1E-160
		AAH36107.1	AAH36107.1 Unknown (protein for MGC:33922)	567	1E-160
		NP_006588.1	NP_006588.1 heat shock 70kDa protein 8 isoform 1; heat shock cognate protein, 71-kDa; heat shock 70kd	565	1E-160
			protein 10; heat shock cognate protein 54; constitutive heat shock protein 70; lipopolysaccharide-		
			associated protein 1; LPS-associated protein 1		
		AAH07276.1	Similar to heat shock cognate 71-kd protein	595	1E-160
		AAD11466.1	AAD11466.1 lteat shock protein	564	564 1E-159
		AAH35665.1	AAH35665.1 heat shock 70kDa protein 6 (HSP70B')	555	1E-157
		NP_002146.1	NP_002146.1 heat shock 70kDa protein 6 (HSP70B'); heat shock 70kD protein 6 (HSP70B'); Heat-shock 70kD protein 6 (HSP70B');	552	1E-156
U 8 9 4 1 5 NULL AAC36522.1	F:2.73 (5to19)	NP_001952.1	NP_001952.1 eukaryotic translation elongation factor 2; polypeptidyl-tRNA	444	1E-125
		XP_170567.1	XP_170567.1 similar to Elongation factor 2 (EF-2)	438	1E-123
	$\neg \neg$				
NM 009242 Mm.35439	39 F:2.73 (5to19)	NP 003109.1	NP 003109.1 secreted protein, acidic, cysteine-rich (osteonectin); Osteonectin (secreted protein, acidic,	575	1E-163

. 2

1E-163 1E-140 1E-133 SE-87 5E-87 5E-87 2E-84 1E-82 4E-60 573 496 474 1028 1028 320 305 229 1030 320 320 6-phosphofructo-2-kinase (EC 2.7.1.105) / fructose-2, 6-bisphosphate 2-phosphatase (EC SPARC precursor (Secreted protein acidic and rich in cysteine) (Osteonectin) (ON) (Basement SPARC precursor (Secreted protein acidic and rich in cysteine) (Osteonectin) (ON) (Basement Chain , Extracellular Matrix Protein Mol id: 1; Molecule: Sparc; Chain: Null; Fragment. Yes; Heterogen: 2 Ca 2+ Ions, One Unidentified Metal Ion Modeled As Ca 2+; Other_details: Carboxy-Terminal Domain (Residues 136 - 286); Synonym: Bm-40, Osteonectin; Engineered. SPARC-like protein 1 precursor (High endothelial venule protein) (Hevin) (MAST 9) Crystallized From 0.7 M K, Na-Tartrate, Ph 7.5 + 2 Mm Cacl2 NP 004557.1 6-phosphofructo-2-kinase/fructose-2,6-biphosphatase 3 AAB99795.1 6-phosphofructo-2-kinase/fructose-2,6-bisphosphatase XP_085281.2 similar to RIKEN cDNA 1700095F04 AAH33721.1 Unknown (protein for MGC:45264) NP_004675.2 SPARC-like 1; mast9; hevin membrane protein BM-40) membrane protein BM-40) unnamed protein product 156 cysteine-rich) osteonectin BAC04065.1 AAA60993.1 pdb|1BMO pdb|1SRA pdb/1NUB Q14515 JC4626 F:2.69 (5to7) NM 026104 Mm.148837 F:2.72 (5to7) A F 2 9 4 6 1 7 Mm. 19669 NP 033268.1 NP 080380.1 AAG02118.1

2

10

15

1005

inducible 6-phosphofructo-2-kinase/fructose 2,6-bisphosphatase

AAC62000.1

	0	0	0	0	P	0	0	0	1E-167	0	0	0	0	0	0
	669	697	889	089	029	699	899	899	589	888	882	882	880	088	871
157	AA06605.1 6-phosphofructo-2-kinase	6-phosphofructo-2-kinase/fructose-2,6-biphosphatase 2 (6PF-2-K/Fru-2,6-P2ASE heart-type isozynie) (PFK-2/FBPase-2) [Includes: 6-phosphofructo-2-kinase; Fructose-2,6-bisphosphatase]	NP_006203.1 6-phosphofructo-2-kinase/fructose-2,6-biphosphatase 2; Fructose-2,6-bisphosphatase, cardiac isozyme	BAB19681.1 6-phosphofructo-2-kinase heart isoform	NP_004558.1 6-phosphofructo-2-kinase/fructose-2,6-biphosphatase 4	6-phosphofructo-2-kinase (EC 2.7.1.105) / fructose-2, 6-bisphosphate 2-phosphatase (EC 3.1.3.46)	NP_002616.1 6-phosphofructo-2-kmase/fructose-2,6-biphosphatase 1; Fructose-2,6-bisphosphatase	6-phosphofructo-2-kinase/fructose-2,6-biphosphatase 1 (6PF-2-K/Fru-2,6-P2ASE liver isozyme Includes: 6-phosphofructo-2-kinase; Fructose-2,6-bisphosphatase 1	6-phosphofructo-2-kinase	Unknown (protein for IMAGE:4712175)	NP_005304.3 glucose regulated protein, 58kDa; glucose regulated protein, 58kD	protein disulfide-isomerase (EC 5.3.4.1) ER60 precursor	ER-60 protein	protein disulfide-isomerase (EC 5.3.4.1) ER60 precursor	BAA03759.1 phospholipase C-alpha
	CAA06605.1	060825	NP_006203.1	BAB19681.1	NP_004558.1	JC5871	NP_002616.1	P16118	CAB06077.1	AAH36000.1	NP_005304.3	JC5704	AAC51518.1	855507	BAA03759.1
										F:2.65 (5to19)			·		
										Mın.709					
							·			NM_007952 Mm.709 NP_031978.1					

	0	4E-92	6E-65	-		4E-64	1E-63	4E-86	1E-85	5E-62	2E-61	3E-57	1E-56	0	0	1E-170	1E-120	1E-103
	198	340	250			247	245	316	314	236	234	219	218	838	983	602	436	380
158	protein disulfide-isomerase (EC 5.3.4.1) ER60 precursor	protein disulfide isomerase related protein (calcium-binding protein, intestinal-related)	NP_000909.2 procollagen-proline, 2-oxoglutarate 4-dioxygenase (proline 4-hydroxylase), beta polypeptide	(protein disulfide isomerase; thyroid hormone binding protein p55); v-erb-a avian erythroblastic	leukemia viral oncogene homolog 2-like	prolyl 4-hydroxylase beta subunit (AA 1-491)	AAA61169.1 thyroid hormone binding protein precursor	NP_066924.1 claudin 1; senescence-associated epithelial membrane protein 1	claudin-1	claudin 7	claudin 7; Clostridium perfringens enterotoxin receptor-like 2; claudin 9	XP_086547.1 similar to Claudin-19	l claudin 19	NP_000228.1 lipoprotein lipase precursor	AAH11353.1 Similar to lipoprotein lipase	AAC61679.1 lipoprotein lipase precursor	NP_06024.1 endothelial lipase precursor; endothelial cell-derived lipase	NP_000227.1 lipase C precursor
	S63994	NP_004902.	NP_000909.			CAA28775.1	AAA61169.	NP_066924.	AAD22962.1	AAH01055.1	NP_001298.	XP_086547.	NP_683763.1 claudin 19	NP_000228	AAH11353.	AAC61679.	NP_006024	NP_000227
								F:2.65 (5to19)						F:2.64 (5to19)				
								NM_016674 Mm.33669 NP 057883.1						NM_008509 Mm.1514 NP_032535.1				

Ŋ

			159		
	ţ	AAA59520.1	A 59520.1 hepatic lipase precursor	379	1E-103
		A28997	triacylglycerol lipase (EC 3.1.1.3) precursor, hepatic	379	1E-103
NM_009690 Mm.6676 F:2.6	F:2.63 (7to19)	NP_005885.1	005885.1 CD5 antigen-like (scavenger receptor cysteine rich family); Spalpha	517	1E-146
NP 033820.1					
		NP_015568.1	015568.1 deleted in malignant brain tumors 1 isoform b precursor	777	4E-74
		CAB56155.1	DMBT1/8kb.2 protein	276	1E-73
		CAC44122.1	DMBT1/8kb.2 protein	275	2E-73
		NP_060049.1	060049.1 deleted in malignant brain tumors 1 isoform c precursor	272	1E-72
		CAB63941.1	DMBT1 prototype	269	1E-71
		BAA78577.1	DMBT1	266	6E-71
		NP_004397.1	004397.1 deleted in malignant brain tumors 1 isoform a precursor	263	5E-70
		836077	M130 antigen	254	3E-67
		138006	M130 antigen precursor, splice form 1	254	3E-67
		138004	M130 antigen precursor, splice form 3	254	3E-67
·		NP_004235.2	NP_004235.2 CD163 antigen; macrophage-associated antigen	254	3E-67
		AAF91396.1	scavenger receptor cysteine-rich type 1 protein M160 precursor	252	1E-66
		NP_542782.1	NP_542782.1 scavenger receptor cysteine rich domain containing, group B (4 domains); scavenger receptor	246	6E-65
			cysteine-rich protein SRCRB-S4D		
		NP_003610.1	003610.1 neurotrypsin precursor; protease, serine, 12; motopsin; brain-specific serine protease 3; leydin	226	9E-59
		AAH07761.1	protease, serine, 12 (neurotrypsin, motopsin)	208	3E-53
NM 025459 Mm.25311 F:2.	F:2.63 (7to19)	BAB15241.1	lunnamed protein product	572	1E-163
				İ	

				Λ9T		
NP_079735.1						
			NP_061873.2	_061873.2 hypothetical protein FLJ20152	571	1E-162
			BAA90982.1	unnamed protein product	204	3E-52
			BAB15252.1	BAB15252.1 unnamed protein product	202	7E-52
X00496	Mm.7043	F:2.63 (5to19)	NP_004346.1	NP_004346.1 hypothetical protein FLJ13902	526	4E-59
CAA25191.1					ì	5
			HLHUG	class II histocompatibility antigen-associated gamma chain	226	4E-59
			CAA27047.1	gamma chain	225	9E-59
			P04233	HLA class II histocompatibility antigen, gamma chain (HLA-DR antigens associated invariant	207	1E-53
				chain) (Ia antigen-associated invariant chain) (Ii) (p33) (CD74 antigen)		
NM_011435 Mm.2407	Mm.2407	F:2.61 (5to19)	NP_003093.1	NP_003093.1 superoxide dismutase 3, extracellular	288	6E-78
NP 035565.1						
NM_007574 Mm.3453	Mm.3453	F:2.56 (5to19)	стилос	complement subcomponent C1q chain C precursor	270	3E-71
NP 031600.1						
			XP_031238.1	XP_031238.1 similar to pancreatic elastase (EC 3.4.21.36) IIIA precursor	269	4E-71
-			BAB71575.1	BAB71575.1 unnamed protein product	268	1E-70
		-				
AK004387		F:2.55 (YtoO)	NP_036276	dynein, axonemal, intermediate polypeptide 1; dynein, axonemal, intermediate chain 1; dynein	171	6E-43
BAC25081.1				intermediate chain DNAI1		
			AAF19816	dynein intermediate chain DNAI1	171	6E-43
			AAH30583	dynein, axonemal, intermediate polypeptide, 1	171	6E-43
NM 008330	008330 Mm.24769	F:2.55 (5to19)	NP 062558.1	NP 062558.1 hypothetical protein R30953 1	228	1E-58

Ŋ

0

20

		558 1E-159		550 1E-156	548 1E-156	510 1E-144	396 1E-110	347 3E-95	266 7E-71	247 3E-65	228 2E-59	roup (nonhistone 324 3E-88		321 3E-87	318 2E-86	310 7E-84	301 2E-81	thle pseudogene; 285 1E-76		otein 2 283 1E-75	283 IE-75	n 1-like 1 250 7E-66	rotein 4 244 4E-64	. 237 6E-62		237 6E-62
161		Unknown (protein for IMAGE:5165618)		calponin 1, basic, smooth muscle; calponins, basic; Calponin 1	smooth muscle cell calponin	h1-calponin	calponin 3; calponin, acidic	calponin 2; Calonin 2	similar to calponin 2; Calonin 2	similar to calponin 2; h2-calponin [Mus musculus]	h2-calponin	NP_002119.1 high-mobility group box 1; high mobility group box 1; high-mobility group (nonhistone	chromosomal) protein 1	HMG-1	nonhistone chromosomal protein HMG-1	dJ579F20.1 (high-mobility group (nonhistone chromosomal) protein 1-like 1)	High mobility group protein 1-like 10 (HMG-1L10)	AAF19244.1 similar to nonhistone chromosomal protein HMG-1 [Homo sapiens]; probable pseudogene;	similar to P09429 (PID:g123369)	high-mobility group box 2; high-mobility group (nonhistone chromosomal) protein 2	high-mobility group (nonhistone chromosomal) protein 2	similar to dJ579F20.1 (high-mobility group (nonhistone chromosomal) protein 1-like 1	NP_005333.1 high-mobility group box 3; high-mobility group (nonhistone chromosomal) protein 4	nuclear antigen Sp100		nııclear autoantigen
		AAH36307		NP 001290	G02142	BAA12983	NP 001830	NP 004359	XP 167021	XP 070819	BAA20887	NP_002119.1		BAA09924.1	S29857	CAB92731.1	9ADA6d	AAF19244.1		NP_002120.1	AAH00903.2	XP_086648.2	NP_005333.1	NP_003104.2	-	AAL77438.1
		F:2.54 (YtoO)										F:2.52 (5to19)														
																	-							-	_	
	NP_032087.1	NM_009922 Mm.4356	NP 034052.1									NM_010439 Mm.16421	NP 034569.1													

			XP_016076.2	016076.2 similar to high mobility group 1	234	3E-61
			S30221	nonhistone chromosomal protein HMG-2B - human	233	9E-61
			XP_116482.1	116482.1 similar to High mobility group protein 4 (HMG-4) (High mobility group protein 2a) (HMG-2a)	212	2E-54
		,	XP_063129.1	063129.1 similar to high mobility group 1	212	2E-54
			XP_115897.1	similar to HMG2a (high mobility group protein 2a)	206	1E-52
			CAA22428.1	HMG2a (high mobility group protein 2a)	206	1E-52
			XP_089930.5	similar to high mobility group 1 protein	205	3E-52
X12905	Mm.8308	F:2.51 (5to19)	NP_002612.1	002612.1 properdin P factor, complement	771	0
CAA31389.1						
			AAB62886.1	properdin	770	0
			\$16150	properdin precursor - human.	768	0
			T45112	properdin [imported] - human	992	0
			T45113	properdin [imported] - human	764	0
			CAA15658.1	dJ212G6.2 (properdin)	731	0
			AAB63279.1	properdin	241	2E-63
			CAC37630.1	fibulin-6	236	7E-62
			AAK68690.1 hemicentin	hemicentin	235	1E-61
			XP_053531.6	053531.6 similar to hemicentin	235	1E-61
			A05319	properdin - human (fragments).	196	7E-50
A K 0 0 7 3 9 2 Mm. 2131	Mm.2131	F:2.49 (5to11)	S70439	pancreatic clastase I (allele HEL1-16) probable splice form I	426	1E-118

l	•	,	ì
ţ	٥)	ľ
			ı
•		7	J

-				163		
BAB25008.1						
			NP_001962.2	001962.2 elastase 1, pancreatic	419	1E-116
			NP_254275.1	254275.1 similar to elastase 1, pancreatic (H. sapiens)	278	7E-74
			BAA00165.1	pancreatic elastase 2 precursor	275	1E-72
			NP_056933.1	pancreatic elastase IIB	268	7E-71
			CAC42422.1	bA265F14.3 (Elastase 2B)	566	5E-70
			NP_031378.1	elastase 3B	253	3E-66
			P08861	Elastase IIIB precursor (Protease E)	253	3E-66
			A29934	pancreatic clastase (EC 3.4.21.36) IIIA precursor	253	4E-66
			NP_005738.2	NP_005738.2 clastase 3, pancreatic (protease E)	252	6E-66
			P09093	Elastase IIIA precursor (Protease E)	252	6E-66
			AAA66350.1	elastase III A	251	2E-65
			AAH05918.1	Similar to elastase 3, pancreatic (protease E)	250	2E-65
			AAA36482.1	protease E precursor	250	3E-65
			S68826	pancreatic elastase (BC 3.4.21.36) isoform 2 precursor	241	1E-62
			Q99895	Caldecrin precursor (Chymotrypsin C)	241	1E-62
			NP_009203.1	_009203.1 chymotrypsin C (caldecrin); caldecrin (serum calcium decreasing factor, elastase IV)	240	2E-62
			CAA74031.1 chymotrypsin	chymotrypsin	236	3E-61
NM_016847 Mm.4351		F:2.48 (5to19)	NP_000697.1	NP_000697.1 arginine vasopressin receptor 1A; V1a vasopressin receptor; vascular/hepatic-type arginine	701	0
NP 058543.1	-			vasopressin receptor; antidiuretic hormone receptor 1A		

Ŋ

opressiu V1B receptor; 1 n receptor n receptor n receptor ssin receptor: ISOTYPE sssin receptor type 2 ssin receptor type 2 pidin pin 1 pin 1 pin 1 pin-3 pin-3 pin-3	P_000907.1 oxytocin receptor 808301A oxytocin receptor RA56562.1, oxytocin receptor P_000045.1 arginine vasopressin receptor 2 913493A vasopressin receptor type 2 AB87678.1 vasopressin receptor type 2 AC08215.1 mucolipidin P_065394.1 mucolipidin AC08215.1 mucolipidin AC07813.1 mucolipidin P_060768.7 nucolipin-3 P_060768.7 proxisonal carnitine octanoyltrans PUKG9 Peroxisonal carnitine octanoyltrans	NP_000907.1 Oxytocin receptor	E B S B I I I E I	000698.1 arginine vasopressin receptor 1B; arginine vasopressin receptor 3; antidiuretic hormone receptor 364 4E-99	oituitary vasopressin receptor 3	355 2E-96	355 2E-96	299 1E-79	244 SE-63	=V2 241 3E-62	216 1E-54	1069 0	1069 0	0 866	0 629	628 1E-179	. 515 IB-145	ransferase (COT)		(EC 2.3.1.137)
	07.1. 4. 4. 4. 4. 4. 1. 15.1. 15.1. 15.1. 13.1. 13.1. 13.1. 13.1.	NP_000907.1 1808301A CAA56562.1, NP_000045.1 1913493A AAB87678.1 CAC08215.1 AAG42242.1 CAC07813.1 AAL84622.1 NP_060768.7 NP_060768.7	Mm.8356 F:2.47 (7to19) CAC08215.1 Mm.8356 F:2.47 (7to19) CAC08215.1 AAG42242.1 CAC07813.1 CAC07813.1 AAL84622.1 AM.28197 F:2.47 (7to19) Q9UKG9	1B; vasopressin V1B receptor; pituitary vasopressin receptor 3		a receptor	n receptor	n receptor	vasopressin receptor 2	ssin receptor:ISOTYPE=V2	ssin receptor type 2	idin	oin 1; mucolipidin	oin 1	oidin	oin-3	in-3	Peroxisonal camitine octanoyltransferase (COT)	camitine O-octanoyltransferase (EC 2.3.1.137)	NP 066974.1 camitine O-octanovltransferase

വ

10

1

 \sim

	٠.			165		
			NP_000746.2	NP_000746.2 carmitine acetyltransferase precursor, isoform 1	305	3E-82
-			NP_003994.2	camitine acetyltransferase isoform 2	305	3E-82
			P43155	Camitine O-acetyltransferase (Camitine acetylase) (CAT).	302	2E-81
A K 0 0 7 5 8 8 NULL XP 135065.1	TULL	F:2.47 (5to7)	AAD30147.1	RING finger protein	214	9E-55
			NP_055060.1	NP_055060.1 ring finger protein 7; sensitive to apoptosis gene	214	1E-54
NM_019985 Mm.30700 NP 064369.1	4m.30700	F:2.46 (7to19)	NP_057593.1	NP_057593.1 C-type lectin-like receptor-2	303	2E-82
·			AAH29554.1	Similar to C-type lectin-like receptor-2	248	6E-66
NM_007572 Mm.370 NP_031598_1	<i>A</i> m.370	F:2.45 (5to19)	NP_057075.1	NP_057075.1 complement component 1, q subcomponent, alpha polypeptide precursor; complement C1q A	286	2E-76
				The state of the s		
AF218416		F:2.44 (7to19)	NP_000361.1	NP_000361.1 tocopherol (alpha) transfer protein (ataxia (Friedreich-like) with vitamin E deficiency);	451	1E-126
AAF25956.1				Tocopherol (alpha) transfer protein		
			G01727	alpha-tocopherol transfer protein	449	1E-125
NM_019642 Mm.22130	Mm.22130	F:2.41 (YtoM)	P04844	lichyl-diphosphooligosaccharideprotein glycosyltransferase 63 kDa subunit precursor	1033	0
NP 062616.1				(Ribophorin II) (RPN-II) (RIBIIR)		
			B26168	ribophorin II precursor	1032	0
			NP_002942.1	P_002942.1 tibophorin II	1028	0
			AAH02380.1	AAH02380.1 Unknown (protein for IMAGE:2961244)	801	0

വ

			. 166	i	
		AAH13028,1	AAH13028,1 Unknown (protein for IMAGE:3532152)	480	IE-135
		CAC34517.1	dJ343K2.2.3 (ribophorin II (isoform 3))	427	1E-119
NM_010442 Mm.17980 NP_034572.1	F.2.41 (71019)	AAH01491.1	heme oxygenase (decycling)	483	1E-136
		NP_002124.1	2 002124.1 heme oxygenase (decyclizing) 1	479	1E-135
		pdb 1QQ8	Chain A, X-Ray Crystal Structure Of Human Heme Oxygenase-1	375	1E-104
		NP_002125.3	_002125.3 heme oxygenase (decyclizing) 2	261	2E-69
		AAB22110.2	heme oxygenase-2; НО-2	255	1E-67
		S21700	heme oxygenase (decyclizing) (EC 1.14.99.3) 2	254	3E-67
		AAA50403.1	AAA50403.1 heme oxygenase	197	3E-50
NM_007833 Mm.1987 NP 0318591	F:2.41 (7to19)	NP_001911.1	NP_001911.1 decorin isoform a preproprotein; dermatan sulphate proteoglycans II; bone proteoglycan II;	584	1E-156
		AAA52301.1 decorin	decorin	554	1E-156
		NP_001702.1	NP_001702.1 biglycan preproprotein; bone/cartilage protcoglycan-I; dermatan sulphate proteoglycan I	382	1E-105
		AAA52287,1 biglycan	biglycan	366	1E-100
		NP_060150.2	NP_060150.2 asporin (LRR class 1); periodontal ligament associated protein 1	352	7E-96
		Q9BXN1	Asportn precursor (Periodontal ligament associated protein-1) (PLAP-1)	352	7E-96
		BAC04007.1	unnamed protein product	314	2E-84
		NP_598011.1	NP_598011.1 decorin isoform b precursor; dermatan sulphate proteoglycans II; bone proteoglycan II;	303	5Ę-81
		NP_598012.1	proteoglycan core protein NP_598012.1 decorin isoform c precursor; dermatan sulphate proteoglycans II; bone proteoglycan II;	238	2E-61
			proteoglycan core protein		

WO 2005/000335 PCT/US2004/017322 (

	2E-51		1E-143	1E-143	1E-143	1E-142	6E-68	9E-78	9E-78	2E-77	2E-74		1E-162	1E-161	2E-77	2E-76	1E-75	4E-56
	204	-	909	505	505	503	256	288	288	287	277		574	570	292	288	285	220
167	BAA90967.1 unnamed protein product		NP_004020.1 interferon regulatory factor 7 isoform b	NP_001563.2 interferon regulatory factor 7 isoform a	Q92985 Interferon regulatory factor 7 (IRF-7)	l interferon regulatory factor 7 isoform d	putative interferon regulatory factor 7C.2	NP_000482.2 complement component 1, q subcomponent, beta polypeptide precursor; complement component C1q, B chain	complement subcomponent C1q chain B precursor [validated	Complement C1q subcomponent, B chain precursor	C1q B-chain precursor		l lumican	lumican	Similar to fibromodulin	Fibromodulin precursor (FM) (Collagen-binding 59 kDa protein) (Keratan sulfate proteoglycan fibromodulin) (KSPG fibromodulin)	NP_002014.1 fibromodulin precursor	NP_008966.1 keratocan; comea plana 2 (autosomal recessive)
	BAA90967.1		NP_004020.	NP_001563.	Q92985	NP_004022.	AAB80691.1	NP_000482.	СІНПОВ	P02746	CAA26880.1		NP_002336.1 lumican	AAA85268.1	AAH35281.1	Q06828	NP_002014.	NP_008966.
			F:2.41 (5to19)					F:2.41 (5to19)				,	F:2.41 (5to19)					
			Mm.3233					Mm.2570										
			NM_016850 Mm.3233 NP_058546.1		ſ			NM_009777 Mm.2570 NP_033907.1					NM_008524 NP_032550.1		·			·

20 Ŋ

		Nm 0003141	108	010	20.00
		141 002 110.1	187 _002/10.1 profitie arguine-fich end feuenie-fich tepeat profess	718	CE-37
		NP_005005.1 osteomodulin	osteomodulin	211	3E-53
NM_010789 Mm.4734	F:2.4 (5to 19)	AAF81641.1	TALE homeobox protein Meis2d	749	0
NF 054919.1		014770	Homeobox protein Meis2 (Meis1-related protein 1)	743	0
		NP_002389.1	NP_002389.1 Meis1 homolog; Meis1 (mouse) homolog	701	0
		AAF81639.1	TALE homeobox protein Meis2b	598	1E-170
		AAF81638.1	TALE homeobox protein Meis2a	591	1E-168
		iAAH01516.1	iAAH01516.1 Unknown (protein for MGC:2820)	570	1E-162
		NP_064534.1	NP_064534.1 homeobox protein Meis2 isoform e; Meis (mouse) homolog 2; Meis1-related gene 1; TALE homeobox protein Meis2	456	1E-128
		789669	Homeobox protein Meis3 (Meis1-related protein 2)	431	1E-120
		AAH25404.1	AAH25404.1 Similar to hypothetical protein DKFZp547H236	409	1E-113
		XP_085929.5	similar to Homeobox protein Meis3 (Meis1-related protein 2)	377	1E-104
		AAM09846.1	AAM09846.1 MEIS1-related protein 2	305	2E-82
NM_013485 Mm.29095 NP 038513.1	F:2.38 (5to19)	CAA69849.1	CAA69849.1 complement 9	599	0
		NP_001728.1	NP_001728.1 complement component 9	999	0
		СЭНО	complement C9 precursor [validated]	629	0
		AAA51889.1	AAA51889.1 C9 complement protein	652	0

2

<u>-</u>1

 $\tilde{\mathbf{z}}$

WO 2005/000335 PCT/US2004/017322

			100		
		NP_000553.1	NP_000553.1 complement component 8, alpha polypeptide precursor	264	4E-70
		C8HUA	complement C8 alpha chain precursor [validated]	263	8E-70
NM_016906 Mm.28375 F:2.37	F:2.37 (5to19)	NP_037468.1	NP_037468.1 Sec61 alpha form 1; sec61 homolog	931	0
NP 058602.1					
		Q9Y2R3	Protein transport protein Sec61 alpha subunit isoform 2 (Sec61 alpha-2)	606	0
		NP_060614.2	NP_060614.2 sec61 homolog; Sec61 alpha form 2	891	0
		AAH02951.1	Similar to CG9539 gene product	828	0
		AAH26179.1	Similar to Sec61 alpha form 2	778	0
		BAB14148.1	BAB14148.1 unnamed protein product	775	0
		BAC11298.1	unnamed protein	969	0
		BAA91692.1	unnamed protein product	432	1E-121
		CAD38592.1	hypothetical protein	425	1E-119
		BAC11283.1	unnamed protein	338	2E-92
A K 0 0 4 9 7 9 Mm.33881 F.2.37 BAB23715.1	F:2.37 (5to19)	XP_091549.1	XP_091549.1 similar to RIKEN cDNA 1300010M03	801	0
		BAB15241.1	BAB15241.1 unuamed protein product	236	2E-60
		BAC11332.1	unnamed protein product	226	1E-57
		NP_061873.2	NP_061873.2 hypothetical protein FLJ20152	208	3E-52

WO 2005/000335

170

Ŋ

NM 013022 Mm 103674 F-2-36 (71019)		XP 094539 s	similar to zinc finger protein KID3	1145	0
NP 038950.1					
		AAM69676 z	zinc finger protein KID3	1125	0
		69	hypothetical protein DKFZp434G1930.1	669	0
NM_009369 Mm.14455 F:2	F:2.36 (5to11)	NP_000349.1	000349.1 transforming growth factor, beta-induced, 68kDa; comeal dystrophy; kerato-epithelin;	1144	0
NP 033395.1			transforming growth factor, beta-induced, 68kD		
		AAC08449.1 E	BIGH3	905	6
		AAN17733.1 e	extracellular matrix protein periostm-bm	571	1E-162
		NP_006466.1	006466.1 osteoblast specific factor 2 (fasciclin I-like); periostin	268	1E-161
		836111	osteoblast-specific factor 2	268	1E-161
		AAC24944.1 BIGH3	BIGH3	332	2E-90
NM_022309 Mm.2018 F:2	F:2.36 (5to19)	NP_074036.1	074036.1 core-binding factor, beta subunit, isoform 1; polyomavirus enhancer binding protein 2, beta	344	SE-94
NP 071704.1			subunit; SL3-3 enhancer factor 1 beta subunit; SL3/AKV core-binding factor beta subunit		
		NP_001746.1	001746.1 core-binding factor, beta subunit, isoform 2; polyomavirus enhancer binding protein 2, beta	302	2E-81
			subunit; SL3-3 enhancer factor 1 beta subunit; SL3/AKV core-binding factor beta subunit		
		6 <i>L</i> 5651	transcription factor CBF beta - human	287	5E-77
		pdb/ICL3	Chain A, Molecular Insights Into Pebp2CBF-Smmhc Associated Acute Leukemia Revealed From	287	5E-77
			The Three-Dimensional Structure Of Pebp2CBF BETA		
		pdb/1H9D	Chain A, Am11CBF-BetaDNA COMPLEX	278	4E-74
8 5 Mm.97986	F:2.35 (5to19)	NP_079282.1	079282.1 hypothetical protein FLJ13373	543	1E-153
BAB31292.1					
		CAD39164.1	CAD39164.1 hypothetical protein	543	1E-153

_	1	
_	_	
	1	

56671.2 methyl-CpG binding protein 1 779 56671.2 methyl-CpG binding domain protein 1 isoform 1 773 51442.1 methyl-CpG binding protein splice variant 1 773 51443.1 methyl-CpG binding protein splice variant 2 662 51443.1 methyl-CpG binding protein splice variant 2 657 51443.1 methyl-CpG binding domain protein 1 isoform PCM1. 632 71735.1 methyl-CpG binding domain protein 1 isoform 3 596 71735.1 methyl-CpG binding domain protein 1 isoform 3 596 71735.1 methyl-CpG binding domain protein 1 isoform 4 526 71735.1 methyl-CpG binding domain protein 1 isoform 4 526 71735.1 methyl-CpG binding domain protein 1 isoform 4 526 71735.1 methyl-CpG binding domain protein 1 isoform 4 526 71735.1 methyl-CpG binding domain protein 1 isoform 3 476 71735.1 methyl-CpG binding domain protein 1 isoform 3 476 71735.1 methyl-CpG binding domain protein 1 isoform 3, precursor, collagen IV, alpha-5 polypeptide, collagen of 476 112-11 71735.1 methyl-CpG binding domain protein 1 isoform 3, precursor, collagen IV, alpha-5 polypeptide, collagen binding bin	•			171	925	ſ
NP_056671.2 methyl-CpG binding protein splice variant 1 773		F:2.35 (5to19)	AAD50371.1	methyl-CpG binding protein 1	6)/	5
NP 056070.2 methyl-CpG binding protein splice variant 1 NP 056070.2 methyl-CpG binding domain protein 1 isoform 2 6572	22.1		NP_056671.2	methyl-CpG binding domain protein 1 isoform 1	779	0
NP_056702 methyl-CpG binding domain protein 1 isoform 2 657 AAD51443.1 methyl-CpG binding domain protein 1 isoform PCM1 NP_056723.2 methyl-CpG binding domain protein 1 isoform PCM1 CAA71735.1 methyl-CpG binding domain protein 1 AAH33242.1 methyl-CpG binding domain protein 1 AAH33242.1 methyl-CpG binding domain protein 1 NP_056691.1 methyl-CpG binding domain protein 1 NP_056691.1 methyl-CpG binding domain protein 1 NP_0356691.1 methyl-CpG binding domain protein 1 NP_0356691.1 methyl-CpG binding domain protein 1 NP_035691.1 methyl-CpG binding domain protein 1 NP_035691.1 methyl-CpG binding domain protein 1 NP_035691.1 methyl-CpG binding domain protein 1 NP_03692891.1 dA24A23.1 (collagen, type IV, alpha 5 (Alport syndrome) NP_0369891.1 methyl-CpG binding domain protein 1 NP_0369891.1 methyl-CpG binding dom			AAD51442.1	methyl-CpG binding protein splice variant 1	773	0
AAD51443.1 methyl-CpG binding protein splice variant 2 NP_056723.2 methyl-CpG binding domain protein 1 isoform PCM1 632 625 626			NP_056670.2	methyl-CpG binding domain protein 1 isoform 2	799	0
NP_056723.2 methyl-CpG binding domain protein 1 isoform PCM1 CAA71735.1 methyl-CpG binding domain protein 1 isoform 1 isoform 3 S97		,	AAD51443.1	methyl-CpG binding protein splice variant 2	657	0
CAA71735.1 methyl-CpG binding domain protein 1 isoform 3 S97			56723.2	methyl-CpG binding domain protein 1 isoform PCM1	632	1E-179
AAH33242.1 methyl-CpG binding domain protein 1 isoform 3 596 NP_05669.1 methyl-CpG binding domain protein 1 isoform 3 596 NP_002375.1 methyl-CpG binding domain protein 1 isoform 4 526 AAH12487.1 Unknown (protein for MGC:21089) 1 MAH12487.1 Unknown (protein for MGC:21089) AAH12487.1 Unknown (protein for MGC:21089) 1 AAA515579 F:2.34 (YtoM) CAB90289.1 alpha 5 type IV collagen, type IV, alpha 5 (Alport syndrome)) AAA51558.1 alpha 5 type IV collagen, isoform 2, precursor; collagen IV, alpha-5 polypeptide; collagen of 476 NP_000486.1 alpha 5 type IV collagen AAA49480.1 alpha-5 type IV collagen AAA99480.1 alpha-5 type IV collage			CAA71735.1	methyl-CpG binding protein	625	1E-177
NP_05669.1 methyl-CpG binding domain protein 1 isoform 3 596 NP_002375.1 methyl-CpG binding domain protein 1 isoform 4 526 AAH12487.1 Uaknown (protein for MGC:21089) 263 Mm.155579 F:2.34 (YtoM) CAB90289.1 dA24A23.1 (collagen, type IV, alpha 5 (Alport syndrome)) 476 NP_203699.1 alpha 5 type IV collagen, isoform 2, precursor, collagen IV, alpha-5 polypeptide; collagen of 476 AAA51558.1 alpha-5 type IV collagen isoform 1, precursor; collagen IV, alpha-5 polypeptide; collagen of 476 AAA51558.1 alpha-5 type IV collagen isoform 3, precursor; collagen IV, alpha-5 polypeptide; collagen of 476 NP_203700.1 alpha 5 type IV collagen isoform 3, precursor; collagen IV, alpha-5 polypeptide; collagen of 476 NP_203700.1 alpha-5 type IV collagen isoform 3, precursor; collagen IV, alpha-5 polypeptide; collagen of 476 NP_203700.1 alpha-5 type IV collagen 5 type IV collagen 5 type IV collagen 5 type IV collagen 6 NP_203700.1 alpha 5 type IV collagen 6 NP_203700.1 alpha 5 type IV collagen 7 type IV collagen 9 NP_203700.1 alpha 5 type IV collagen 1 type IV colla				methyl-CpG binding domain protein 1	597	1E-169
Mm.155579 Fr.2.34 (YroM) CAB90289.1 dA24A23.1 (collagen, type IV, alpha 5 (Alport syndrome)) Mm.155579 Fr.2.34 (YroM) CAB90289.1 dA24A23.1 (collagen, type IV, alpha 5 (Alport syndrome)) AAAA51558.1 alpha 5 type IV collagen, isoform 2, precursor; collagen IV, alpha-5 polypeptide; collagen of 476 basement membrane, alpha-5 chain AAAA51558.1 alpha-5 type IV collagen, basement membrane, alpha-5 chain AAAA99480.1 alpha-5 type IV collagen alpha-5 chain AAAA99480.1 alpha-5 type IV collagen isoform 3, precursor; collagen IV, alpha-5 polypeptide; collagen of 476 basement membrane, alpha-5 chain AAA99480.1 alpha-5 type IV collagen NP_203700.1 alpha 5 type IV collagen, isoform 3, precursor; collagen IV, alpha-5 polypeptide; collagen of 476 basement membrane, alpha-5 chain				methyl-CpG binding domain protein 1 isoform 3	969	
AAH12487.1 Unknown (protein for MGC:21089) Mm.155579 F:2.34 (YtoM) CAB90289.1 dA24A23.1 (collagen, type IV, alpha 5 (Alport syndrome)) NP_203699.1 alpha 5 type IV collagen, isoform 2, precursor; collagen IV, alpha-5 polypeptide; collagen of 476 AAAA51558.1 alpha-5 type IV collagen NP_000486.1 alpha-5 type IV collagen AAA99480.1 alpha-5 type IV collagen AAAA99480.1 alpha-5 type IV collagen, isoform 3, precursor; collagen IV, alpha-5 polypeptide; collagen of 476 NP_203700.1 alpha-5 type IV collagen, isoform 3, precursor; collagen IV, alpha-5 polypeptide; collagen of 476 NP_203700.1 alpha-5 type IV collagen, isoform 3, precursor; collagen IV, alpha-5 polypeptide; collagen of 476		,		methyl-CpG binding domain protein 1 isoform 4	526	
Mm.15579 F:2.34 (YtoM) CAB90289.1 dA24A23.1 (collagen, type IV, alpha 5 (Alport syndrome)) NP_203699.1 alpha 5 type IV collagen, isoform 2, precursor; collagen IV, alpha-5 polypeptide; collagen of 476 AAA51558.1 alpha-5 type IV collagen NP_000486.1 alpha 5 type IV collagen, isoform 1, precursor; collagen IV, alpha-5 polypeptide; collagen of 476 basement membrane, alpha-5 chain AAA99480.1 alpha-5 type IV collagen NP_203700.1 alpha 5 type IV collagen, isoform 3, precursor; collagen IV, alpha-5 polypeptide; collagen of 476 basement membrane, alpha-5 chain			AAH12487.1	Unknown (protein for MGC:21089)	263	
Mm.155579 F.2.34 (YtoM) CAB90289.1 dA24A23.1 (collagen, type IV, alpha 5 (Alport syndrome)) NP_203699.1 alpha 5 type IV collagen, isoform 2, precursor; collagen IV, alpha-5 polypeptide; collagen of 476 AAA51558.1 alpha-5 type IV collagen NP_000486.1 alpha 5 type IV collagen AAA99480.1 alpha-5 type IV collagen AAA99480.1 alpha-5 type IV collagen NP_203700.1 alpha 5 type IV collagen, isoform 3, precursor; collagen IV, alpha-5 polypeptide; collagen of 476 AAA99480.1 alpha-5 type IV collagen NP_203700.1 alpha 5 type IV collagen, isoform 3, precursor; collagen IV, alpha-5 polypeptide; collagen of 476 basement membrane, alpha-5 chain					_	
NP_203699.1 alpha 5 type IV collagen, isoform 2, precursor; collagen IV, alpha-5 polypeptide; collagen of 476 AAA51558.1 alpha-5 type IV collagen NP_000486.1 alpha-5 type IV collagen, isoform 1, precursor; collagen IV, alpha-5 polypeptide; collagen of 476 basement membrane, alpha-5 chain AAA99480.1 alpha-5 type IV collagen NP_203700.1 alpha-5 type IV collagen, isoform 3, precursor; collagen IV, alpha-5 polypeptide; collagen of 476 basement membrane, alpha-5 chain		F:2.34 (YtoM)	CAB90289.1	dA24A23.1 (collagen, type IV, alpha 5 (Alport syndrome))	476	
basement membrane, alpha-5 chain 1000486.1 alpha-5 type IV collagen, isoform 1, precursor; collagen IV, alpha-5 polypeptide; collagen of 476 basement membrane, alpha-5 chain 109480.1 alpha-5 type IV collagen 203700.1 alpha-5 type IV collagen, isoform 3, precursor; collagen IV, alpha-5 polypeptide; collagen of 476 basement membrane, alpha-5 chain			NP_203699.1	alpha 5 type IV collagen, isoform 2, precursor; collagen IV, alpha-5 polypeptide; collagen of	İ	
A 156.1 alpha-5 type IV collagen 1000486.1 alpha 5 type IV collagen, isoform 1, precursor; collagen IV, alpha-5 polypeptide; collagen of 476 basement membrane, alpha-5 chain basement membrane, isoform 3, precursor; collagen IV, alpha-5 polypeptide; collagen of 476 basement membrane, alpha-5 chain				basement membrane, alpha-5 chain		
000486.1 alpha 5 type IV collagen, isoform 1, precursor; collagen IV, alpha-5 polypeptide; collagen of 476 basement membrane, alpha-5 chain collagen 1 alpha-5 type IV collagen collagen, isoform 3, precursor; collagen IV, alpha-5 polypeptide; collagen of 476 basement membrane, alpha-5 chain				alpha-5 type IV collagen	476	
basement membrane, alpha-5 chain 99480.1 alpha-5 type IV collagen 203700.1 alpha 5 type IV collagen, isoform 3, precursor; collagen IV, alpha-5 polypeptide; collagen of 476 basement membrane, alpha-5 chain			NP_000486.1	alpha 5 type IV collagen, isoform 1, precursor; collagen IV, alpha-5 polypeptide; collagen of		
99480.1 alpha-5 type IV collagen 476 203700.1 alpha 5 type IV collagen, isoform 3, precursor; collagen IV, alpha-5 polypeptide; collagen of 476 basement membrane, alpha-5 chain				basement membrane, alpha-5 chain		
203700.1 alpha 5 type IV collagen, isoform 3, precursor; collagen IV,alpha-5 polypeptide; collagen of basement membrane, alpha-5 chain			AAA99480.1	alpha-5 type IV collagen	476	
basement membrane, alpha-5 chain			NP_203700.1	alpha 5 type IV collagen, isoform 3, precursor; collagen IV, alpha-5 polypeptide; collagen of		1E-13
				basement membrane, alpha-5 chain		

		1/2		
	AAA52045.1	collagen type IV alpha 5 chain	476	1E-134
	CAC13153.1	bA472K17.2 (collagen type IV alpha 1)	421	1E-118
	AAA52006.1	pro-alpha-1(IV)	421	1E-118
	AAH47305.1	Similar to collagen, type IV, alpha 1	421	1E-118
	AAA52042.1	procollagen alpha-1 type IV	421	1E-118
	NP_001836.1	NP_001836.1 alpha 1 type IV collagen preproprotein; collagen IV, alpha-1 polypeptide; collagen of basement	421	1E-118
		membrane, alpha-1 chain		
	ILIIA	Chain A, The 1.9-A Crystal Structure Of The Noncollagenous (Nc1)Domain Of Human Placenta	419	1E-117
		Collagen Iv Shows Stabilization Via A Novel Type Of Covalent Met-Lys Cross-Link.		
	1402236A	collagen alpha1(IV)	417	1E-117
	AAM97359.1	arresten	415	1E-116
	NP_000082.1	NP_000082.1 alpha 3 type IV collagen, isoform 1, precursor; collagen IV, alpha-3 polypeptide (goodpasture	365	1E-101
		antigen)		
	CGHU3B	collagen alpha 3(IV) chain precursor, long splice form	365	1E-101
	CAC36101.1	alpha3 type IV collagen	365	1E-101
	AAAS1556.1	alpha-3 type IV collagen	365	1E-101
	AAA21610.1	alpha-3 type IV collagen	365	1E-101
	AAF72632.1	tuntstatin	365	1E-101
	AAB19637.1	type IV collagen alpha 3 chain	351	5E-97
	CAA29098.1	alpha (2) chain	318	6E-87
	P08572	Collagen alpha 2(IV) chain precursor	318	6E-87
	NP 001837.1	alpha 2 type IV collagen preproprotein; canstatin	318	6E-87
	ILIIC	1LIIC Chain C, The 1.9-A Crystal Structure Of The Noncollagenous (Nc1)Domain Of Human Placenta	a 318	6E-87
		Collagen Iv Shows Stabilization Via A Novel Type Of Covalent Met-Lys Cross-Link.		
	AAA52043.1	alpha-2 type IV collagen	317	1E-86
	AAA58422.1	collagen alpha-2 type IV	316	2E-86
	AAF72631.1	canstatin	- 1	
	NP_378667.1	NP_378667.1 type IV alpha 6 collagen, isoform B precursor; collagen IV, alpha-6 polypeptide; collagen of	f 308	4E-84
		basement membrane, alpha-6	_	
	AAB19039.1	collagen type IV a6 chain	308	4E-84
	Q14031	Collagen alpha 6(IV) chain precursor	308	4E-84
	AAB19038.1	AAB19038.1 collagen type IV a6 chain	308	4E-84

വ

			1.73		
		NP_001838.1	001838.1 type IV alpha 6 collagen, isoform A precursor; collagen IV, alpha-6 polypeptide; collagen of	308	4E-84
		·	basement membrane, alpha-6		
		BAA04809.1	collagen	308	4E-84
		NP_000083.1	alpha 4 type IV collagen precursor; Collagen IV, alpha-4 polypeptide; collagen of basement	286	2E-77
			membrane, alpha-4 chain		
		BAA04214.1	alpha 4(IV) collagen	286	2E-77
		B49736	collagen alpha 3(IV) chain, medium splice form - human	233	2E-61
		AAA18942.1	collagen type IV alpha 3 .	233	2E-61
	-	NP_112730.1	NP_112730.1 alpha 3 type IV collagen, isoform 2, precursor; collagen IV, alpha-3 polypeptide (goodpasture	233	2E-61
-			antigen)		
NM_010906 Mm.9394	F:2.34 (YtoM)	NP_002492	nuclear factor I/X (CCAAT-binding transcription factor)	738	0
NP 035036.1					
		AAB52369	nuclear factor I	719	0
		AAB52371	nuclear factor I	692	0
		Q14938.	Nuclear factor 1 X-type (Nuclear factor 1/X) (NF1-X) (NF1-X) (NF-I/X) (CCAAT-box binding	692	0
			transcription factor) (CIF) (TGGCA-binding protein).		
		XP 046827	similar to transcription factor NF1 [Rattus norvegicus]	524	1E-148
		012857	Nuclear factor 1 A-type (Nuclear factor 1/A) (NF1-A) (NF1-A) (NF-I/A) (CCAAT-box binding	524	1E-148
			transcription factor) (CTF) (TGGCA-binding protein).		
		BAA92677	KIAA1439 protein	524	1E-148
		P08651	Nuclear factor 1 C-type (Nuclear factor 1/C) (NF1-C) (NFI-C)(NF-I/C) (CCAAT-box binding	429	1E-120
			transcription factor) (CTF) (TGGCA-binding protein).		
		AAH:12120	nuclear factor I/C (CCAAT-binding transcription factor)	428	1E-120
		NP 005588	nuclear factor I/C (CCAAT-binding transcription factor)	427	1E-119
		B33416.	nuclear factor I	426	1E-119
		S01038	transcription factor, CCAAT-binding	416	1E-116
		AAA93126	nuclear factor 1 X-type	410	1E-114
		AAH01283	Similar to nuclear factor I/B	410	1E-114
		NP 005587		409	1E-114
		AAC15752	NFI-X_HUMAN NUCLEAR FACTOR 1/X (NFI-X); CCAAT-BOX BINDING	390	1E-108
			TRANSCRIPTION FACTOR (CTF), TGGCA-BINDING PROTEIN		

വ

		10000	-	305	16 107
			nuclear factor I.X. [AA 187-441]; nuclear factor I/X; NFIX_HUMAN NUCLEAR FACTOR I/X;	354	2E-97
			NFI-X; NF-IX; TGGCA-BINDING PROTEIN; CTF		
		AAA93125	nuclear factor 1 B-type	352	7E-97
-		\B51197	nuclear factor I B3	350	4E-96
			CTF5	343	3E-94
		AAC32594	CIF-1	331	2E-90
		AAD38241	NFI-X3=transcription factor	307	3E-83
		AAB52370	nuclear factor I	216	6E-56
A K 0 1 1 4 9 5 Mm.203896 F:2.34 (5to19)	7:2.34 (5to19)	NP_005561.1	NP_005561.1 lectin, mannose-binding, 1 precursor; intracellular mannose specific lectin; endoplasmic	849	0
BAB27655.1			reticulum-golgi intermediate compartment protein 53		
	1	S42626	ER-golgi intermediate compartment protein	848	0
·		AAH32330.1	AAH32330.1 lectin, mannose-binding, 1	848	0
					_
		1919261A	protein ERGIC-53	844	0
		NP_068591.1	lectin, mannose-binding, 1 like; BRGL protein; BRGIC-53-like protein	255	2E-66
NM_010354 Mm.21109 F	F:2.34 (5to19)	NP_000168.1	NP_000168.1 gelsolin (amyloidosis, Finnish type); Gelsolin	1422	0
NP 034484.1					
		Q9Y6U3	Adseverin (Scinderin)	904	0
		BAC11416.1	unnamed protein product	904	0
		AAK60494.1	scinderin	668	0
		NP_009058.1	NP_009058.1 villin 1; Villin-1	672	0
		AAD15423.1	similar to mouse adseverin(D5); similar to PID:g2218019	999	0
·		BAB67798.1	KIAA 1905 protein	999	0
		pdb 1DB0	Chain A, Carboxy-Terminal Half Of Gelsolin (G4-G6) Bound To Actin	643	0

1.75

			Τ/2		
		NP_006567.2 advillin	advillin	640	6
			Advillin (p92)	638	0
	,	NP_149119.1	scinderin; adseverin; KIAA1905 protein	588	1E-167
		AAH17491.1	Similar to gelsolin (amyloidosis, Finnish type)	542	1E-153
		BAC11465.1	unnamed protein product	497	1E-139
		AAH04134.1	Similar to advillin	464	1E-129
		pdb 1JHW	Chain A, Ca2+-Binding Mimicry In The Crystal Structure Of The Bu3+- Bound Mutant Human	389	1E-106
			Macrophage Capping Protein Cap G		\top
\neg					
NM_010238 Mm.3444 F.2.3 NP_034368.1	F:2.33 (7to19)	NP_005095.1	005095.1 bromodomain containing protein 2; female sterile homeotic-related gene 1	1083	0
		CAC69991.1	O14.1.1 (bromodomain-containing protein 2 (RING3, KIAA9001), isoform 1)	1082	0
		A56619	female sterile homeotic (fsh) homolog RING3	1048	0
		50.1	kinase	1046	0
		NP_031397.1	031397.1 bromodomain containing protein 3; bromodomain-containing 3; RING3-like gene; open reading frame X	642	0
		AAC27978.1	R31546_1	577	1E-164
		NP_490597.1	490597.1 bromodomain-containing protein 4 isoform long; similar to RING3; chromosome-associated protein	577	1E-164
		NP_055114.1	055114.1 bromodomain-containing protein 4 isoform short; similar to RING3; chromosome-associated protein	577	1E-164
		NP_001717.1	001717.1 testis-specific bromodomain protein	905	1E-142
,		AAH32124.1	Similar to bromodomain containing 3	484	1E-136

2 0 1

Ŋ

 $\ddot{1}$

-1

 \sim

		AAF62895.1	protein arginine N-methyltransferase 1-variant 1	673	0
		AAF62894.1	protein arginine N-methyltransferase 1-variant 3	673	0
		CAA71765.1	arginine methyltransferase	929	0
		CAA71763.1	arginine methyltransferase	029	0
		NP_001527.1	001527.1 HMT1 hnRNP methyltransferase-like 2	637	0
		Q9NR22	Protein arginine N-methyltransferase 4	562	1E-160
		NP_062828.2	062828.2 HMT1 hnRNP methyltransferase-like 3	562	1E-160
		AAH19339.1	Unknown (protein for IMAGE:3027997)	296	4E-80
		060678	Protein arginine N-methyltransferase 3	296	4E-80
-					
A F 3 2 0 9 9 6 Mm.14569 F: AAK73808.1	F:2.32 (7to19)	NP_057712.2	057712.2 WW domain-containing adapter with a coiled-coil region isoform 1	1044	0
		NP_567822.1	567822.1 WW domain-containing adapter with a coiled-coil region, isoform 2	951	0
		BAB71029.1	unnamed protein product	949	0
		AAH04258.1	hypothetical protein PRO1741	938	0
		CAC16000.1	bA48B24.1 (A novel protein containing a formin binding protein (FBP28) domain)	861	0
		CAD28517.1	CAD28517.1 hypothetical protein	288	1E-167
		BAB47473.1	KIAA1844 protein	533	1E-150
		NP_567823.1	567823.1 WW domain-containing adapter with a coiled-coil region, isoform 3	521	1E-146

	178	hypothetical protein MGC10753.	
		AAH10356	
*	:		

		AAH10356	hypothetical protein MGC10753	350	5E.05
		T			_L_
NM_020564 Mm.6562 NP_065589.1	F:2.32 (5to19)	AAC78553.1	hydroxysteroid sulfotransferase SULT2B1a	228	1E-59
		NP_004596.1	004596.1 sulfotransferase family, cytosolic, 2B, member 1; sulfotransferase family 2B, member 1	228	1E-59
		AAC78499.1	hydroxysteroid sulfotransferase SULT2B1b	228	1E-59
		21465697	Chain A, Crystal Structure Of Human Dehydroepiandrosterone Sulfotransferase In Complex With	h 218	1E-56
		r	Substrate		
				_	
NM_007614 Mni.3476	F:2.31 (YtoM)	NP_001895.1	001895.1 catenin (cadherin-associated protein), beta 1, 88kDa; catenin (cadherin-associated protein), beta	a 1523	0
NP 031640.1			1 (88kD); catenin (cadherin-associated protein), beta 1 (88kDa		
		pdb 1JPW	Chain A, Crystal Structure Of A Human Tcf-4 BETA-Catenin Complex	1026	0
		pdb[1G3J	Chain A, Crystal Structure Of The XtcB-CbdBETA-Catenin Armadillo Repeat Complex	1014	0
		pdb 1JDH	Chain A, Crystal Structure Of Beta-Catenin And Htcf-4	1007	0
		·			
		BAB93475.1	catenin beta 1	994	0
·		AAH00441.1	junction plakoglobin	929	0
		NP_002221.1	002221.1 junction plakoglobin, isoform 1; gamma-catenin	929	0
		P14923	Junction plakoglobin (Desmoplakin III	913	0
: -		AAA64895.1	A64895.1 Plakoglobin	912	0
		AAL89457.1	beta-catenin	612	1E-174
		2121362A	plakoglobin	293	1E-77
				_	

IJ

	0	0	0	0	1E-121	1E-121	1E-121	1E-121	2B-62		2E-62	3E-59	1E-58	1E-58	1E-132	1E-128	1E+128
	984	983	931	876	439	439	404	437	241		241	231	229	229	469	456	456
212	AC34298.1 DEAD box RNA helicase DDX3	dead box, X isoform	NP_004651.2 DEAD/H (Asp-Glu-Ala-Asp/His) box polypeptide, Y chromosome; DEAD/H box-3, Y-linked	DEAD-box protein 3, Y-chromosomal	VASA protein	hypothetical protein		BEAD box RNA helicase	Ectonucleoside triphosphate diphosphohydrolase I (NTPDase I) (Ecto-ATP diphosphohydrolase	(ATPDase) (Lymphoid cell activation antigen) (Ecto-apyrase) (CD39 antigen)	NP_001767.2 ectonucleoside triphosphate diphosphohydrolase 1; CD39 antigen	Ectonucleoside triphosphate diphosphohydrolase 2 (NTPDase2) (Ecto-ATPase) (CD39 antigen-like 1)	ectonucleoside triphosphate diphosphobydrolase 3; CD39-like 3	E-type ATPase	cytokine-inducible SH2-containing protein isoform 2; cytokine-inducible SH2-containing protein;	cytokine-inducible SH2-containing protein isoform 1; cytokine-inducible SH2-containing protein; cytokine-inducible inhibitor of signaling type 1B; suppressor of cytokine signaling	cytokine-inducible inhibitor of signalling type 1b
	AAC34298.1	AAC51829.1	NP_004651.2	015523	AAF72705.1	CAB70750.1	XP_066968.2	AAF86585.1	P49961		NP_001767.2	Q9Y5L3	NP_001239.1	AAC09236.2	NP_659508	NP_037456	AAF97410
	F:2.14 (5to19)								F:2.14 (5to19)						F:2.13 (YtoO)		
									Mm.33403		1			,	Mm.4592		
	NM_008015 Mm.18459 NP_032041.1								A K 0 0 8 5 9 0 Mm.33403	BAB25764.1					NM_009895 Mm.4592 NP 034025.1		

			717		
		JC5695	Dum!p/Vps1p-like protein	1256	0
		AAH00136.1	AAH00136.1 Similar to dynamin 1-like	1211	0
	•				
		A40671	dynamin, internal form 1, long C-terminal form	456	1E-126
		B40671	dynamin, internal form 2, short C-terminal form	456	1E-126
		NP_004399.1	NP_004399.1 dynamin 1; dynamin; dynamin1; Dynamin-1	456	1E-126
		JC4305	dynamin II - human	441	1E-122
		P50570	Dynamin 2	440	1E-122
		NP_004936.1	NP_004936.1 dynamin 2; Dynamin II	439	1E-121
		BAA74843.2	KIAA0820 protein	429	1E-118
		XP_044463.5	XP_044463.5 similar to Dynamin 3 (Dynamin, testicular) (T-dynamin)	429	1E-118
		NP_056384.1	NP_056384.1 KIAA0820 protein	428	1E-118
		CAB92724.1	CAB92724.1 bA277C14.1 (novel Dynamin family member (KIAA0820))	324	324 1.5E-86
NM_013562 Mm.168 F	F:2.15 (5to19)	6162421	similar to INTERFERON-RELATED DEVELOPMENTAL REGULATOR 1 (NERVE GROWTH FACTOR-INDITCIBLE PROTEIN PC4)	819	0
		7387801	INTERFERON-RELATED DEVELOPMENTAL REGULATOR 1 (NERVE GROWTH	816	0
			FACTOR-INDUCIBLE PROTEIN PC4)		
		4504607	interferon-related developmental regulator 1	774	0
		AAC24562.1	similar to mouse interferon-related protein PC4; 96% identical to P19182 (PID:g135861)	516	1E-146
		Q12894	Interferon-related developmental regulator 2 (SKIMC15 protein)	409	1E-114
		AAC16924.1	interferon-related putative protein	409	1E-114
		NP_006755.2	NP_006755.2 interferon-related developmental regulator 2; Interferon-related protein	409	1E-114
		AAH01327.1	AAH01327.1 interferon-related developmental regulator 2	407	1E-113

0

H

 $\bar{\mathbf{v}}$

		210		
	CAA23782.1	A23782.1 histocompatibility antigen HLA-DR	214	2E-55
	172480	MHC cell surface glycoprotein	204	1E-52
	154437	MHC class II DQ-alpha protein - human (fragment).	199	4E-51
	168717	gene HLA-DQA2 protein - human (fragment)	197	1E-50
A K 0 0 8 2 7 3 Mm.2241 F:2.15 (7to19)	NP_001166.1	001166.1 Rho GDP dissociation inhibitor (GDI) beta; Ly-GDI	270	3E-71
XP_132918.1				·
	pdb/1DS6	Ras-Related C3 Botulinum Toxin Substrate 2	267	2E-70
	CAA45344.1	CAA45344.1 rho GDP dissociation inhibitor (GDI)	234	1E-60
	NP_004300.1	004300.1 Rho GDP dissociation inhibitor (GDI) alpha	234	2E-60
	pdb 1FST	Chain A, Crystal Structure Of Truncated Human Rhogdi Triple Mutant	231	1E-59
	pdb 1FT0	Chain A, Crystal Structure Of Truncated Human Rhogdi K113a Mutant	226	6E-58
	pdb/1FT3	Chain A, Crystal Structure Of Truncated Rhogdi K141a Mutant	226	6E-58
	pdb 1RHO	Chain A, Structure Of Rho Guanine Nucleotide Dissociation Inhibitor	223	3E-57
	pdb 1FSO	Chain A, Crystal Structure Of Truncated Human Rhogdi Quadruple Mutant	223	3E-57
			-	
A K 0 1 8 1 9 5 Mm.140013 F:2:15 (5to19)	NP_005681.1	. 005681.1 dynamin 1-like protein, isoform 3; dynamin-like protein	1274	0
	AAH24590.1	AAH24590.1 dynamin 1-like	1270	0
	NP_036193.1	036193.1 dynamin 1-like protein, isoform 2; dynamin-like protein	1269	0
	BAA22193.1	Dınn1p/Vps1p-like protein	1260	0
	AAD39541.1	AAD39541.1 dynamin-like protein DYNIV-11	1260	0
	NP_036192.1	036192.1 dynamin 1-like protein, isoform 1; dynamin-like protein	1259	0

	7E-67	2E-65	8E-65	8E-65	8E-65	2E-64	2E-63	8E-62	2E-60	7	2E-57	2E-57		2E-57	2E-57		2E-57		2E-57		2E-57		1E-55
	252	247	245	245	245	244	240	235	231		220	220		220	220		220		220		220		214
209	P_002110.1 major histocompatibility complex, class II, DO alpha; lymphocyte antigen; HLA-D0-alpha; major histocompatibility complex, class II, DN alpha	HLA-DNA1.1a	XP_042473.2 major histocompatibility complex, class II, DR alpha precursor	hla-dr antigen heavy chain (aa 3 at 60)	HLA-DR alpha heavy chain	major histocompatibility complex, class II, DR alpha precursor; HLA class II histocompatibility antigen, DR alpha chain	MHC class II antigen DQ-alpha-1 chain	antigen HLA Dratpha	Chain A, Hla-Dr1 (Dra, Drb1 0101) Human Class Ii Histocompatibility Protein (Extracellular	Domain) Complexed With Endogenous Peptide	Chain A, The Structure Of An Intermediate In Class Ii Mhc Maturation: Clip Bound To Hla-Dr3	Chain A, Crystal Structure Of Hla-Dr2 (Dra0101,Drb11501) Complexed With A Peptide From	Human Myelin Basic Protein	Chain A, Structure Of The Epstein-Barr Virus Gp42 Protein Bound To The Mhc Class Ii	Chain A, Crystal Structure Of A Complex Of A Human AlphaBETA-T Cell Receptor, Influenza	Ha Antigen Peptide, And Mhc Class Ii Molecule, Hla-Dr4	Chain A, Crystal Structúre Of Staphylococcal Enterotoxin H In Complex With Human Mhc Class	II	Chain A, Crystal Structure Of Hla-Dr1TPI(23-37, Thr28>ile Mutant) Complexed With	Staphylococcal Enterotoxin C3 Variant 3b2 (Sec3-3b2)	Chain A, Crystal Structure Of Hla-Dr I TPI(23-37) Complexed With Staphylococcal Enterotoxin	C3 Variant 3b2 (Sec3-3b2)	antigen HLA Daalpha
	NP_002110.	BAA81787.1	XP_042473.	AAA36301.1	CAA25076.1	NP_061984.1	AAB70189.1	1102205B	pdb 1AQD	1	pdb 1A6A	pdb 1BX2		pdb 1KG0	pdb 1J8H		pdb 1HXY		pdbllKLG	·	pdb/1KLU		1102205A
		,																			-		

NP_034508.1				
			_	
	A29312	MHC class II histocompatibility antigen HLA-DQ alpha chain precursor	347	1E-95
	1004300B	antigen alpha,II histocompatibility	346	3E-95
	A37044	MHC class II histocompatibility antigen HLA-DQ alpha 1 (DQw4 specificity) precursor	345	7E-95
	P04226	HLA class II histocompatibility antigen, DQ(2) alpha chain precursor	344	1E-94
,	NP_002113.2	3.2 major histocompatibility complex, class II, DQ alpha 1 precursor	343	2E-94
	AAK11577.1	7.1 MHC class II antigen precursor	343	2E-94
	P01907	HLA class II histocompatibility antigen, DQ(5) alpha chain precursor (DC-1 alpha chain)	341	1E-93
	1004300C	antigen alpha,II histocompatibility	340	1E-93
	AAA52671.1	1 MHC HLA-DQA1-alpha protein	340	2E-93
	P01908	HLA class II histocompatibility antigen, DQ(1) alpha chain precursor (DC-4 alpha chain)	339	3E-93
	NP_06444	064440.1 major histocompatibility complex, class II, DQ alpha 2	338	9E-93
,	AAB51233	AAB51233.1 MHC class II HLA-DQ-alpha chain	337	1E-92
	AAA59834	A59834.1 MHC HLA-DX-alpha chain	337	2E-92
	HLHUDX	MHC class II histocompatibility antigen HLA-DQ alpha 2 chain	337	2E-92
	154290	cell surface glycoprotein - human	333	2E-91
	P05536	HLA class II histocompatibility antigen, DQ(W3) alpha chain precursor	331	8E-91
•	XP_17526	175260.1 similar to HLA class II histocompatibility antigen, DQ(3) alpha chain precursor (DC-alpha)	331	8E-91
		(HLA-DCA) (HLA-DQA1*05011)		
	AAA69564	A69564.1 HLA DQA*0302	325	5E-89
`	AAA59754.1	i.1 HLA-DQ alpha chain	325	5E-89
	P04225	HLA class II histocompatibility antigen, DQ(4) alpha chain precursor (DQ-DRW9 alpha chain)	1) 323	2E-88
	154444		320	2E-87
	1612294A	HLA DQw4 alpha	318	6E-87

Ŋ

0

S

v	
_	
_	
•	
м	
-	

				502		
NM_011415 Mm.4272 NP_035545.1		F:2.16 (5to11)	NP_003059.1	_003059.1 snail 2; neural crest transcription factor SLUG; slug (chicken homolog), zinc finger protein	458	1E-128
			NP_005976.2	NP_005976.2 snail 1 homolog; snail 1 zinc finger protein	248	2E-65
			AAD17332.1	zinc finger protein	248	2E-65
			XP_065615.1	similar to snail 1 (drosophila homolog), zinc finger protein	2111	4E-54
A K 0 1 1 3 0 6 Mm.196607 F:2.16 (5to19) BAB27532.1	n.196607	F:2.16 (5to19)	NP_001961.1	NP_001961.1 eukaryotic translation initiation factor 5A; eIF5AI; eIF5A	311	1E-83
			153801	gene eif-5A protein	302	6E-81
			XP_016093.1	XP_016093.1 similar to eukaryotic initiation factor 5A	300	1E-80
	·		NP_065123.1	NP_065123.1 eIF-5A2 protein; eIF5AII	265	1E-69
NM_007686Mm.117180 F:2.16 (5to19)	n.117180	F:2.16 (5to19)	CAA68416.1 factor I	factor 1	832	0
NP 031712.1			#^			
			P05156*,	Complement factor I precursor (C3B/C4B inactivator)	832	0
			NP_000195.1 *,	NP_000195.1 I factor (complement)	830	0
			AAH20718.1,	AH20718.1, Similar to I factor (complement)	434	1E-120
			CAA68417.1,	CAA68417.1, heavy chain of factor I	422	IE-117
			CAA68418.1,	CAA68418.1, light chain of factor I	414	1E-114
			1202205A	complement factor I light chain	328	2E-88
NM 010378 Mn	n.175310	010378 Mm.175310 F:2.15 (11to19)	нгнирс	MHC class II histocompatibility antigen HLA-DC-4 alpha chain precursor	347	1E-95

ហ

7

J U ر ا

			205		
		JN0619	glucuronosyltransferase (EC 2.4.1.17) 2B-4 precursor	702	0
		AAC32272.1	UDP glucuronosyltransferase 2B4 precursor	269	0
		NP_001065.1	NP_001065.1 UDP glycosyltransferase 2 family, polypeptide B7; UDP-glucuronyltransferase, family 2, beta-7	692	0
		S11309	glucuronosyltransferase (EC 2.4.1.17)	691	0
		AAH30974.1	UDP glycosyltransferase 2 family, polypeptide B7	069	0
		NP_066962.1	NP_066962.1 UDP glycosyltransferase 2 family, polypeptide B4; UDP-glucuronyltransferase, family 2, beta-4	889	0
		NP_001064.1	NP_001064.1 UDP glycosyltransferase 2 family, polypeptide B11	677	0
		JE0200	orplian UDP-glucuronosyltransferase (EC 2.4)	229	0
		NP_001066.1	NP_001066.1 UDP glycosyltransferase 2 family, polypeptide B10	099	0
		NP_444267.1	444267.1 UDP glycosyltransferase 2 family, polypeptide B28	099	0
		NP_006789.1	NP_006789.1 UDP glycosyltransferase 2 family, polypeptide A1; UDP glucuronosyltransferase 2 family, polypeptide A1	579	1E-165
					T
NM_015784 Mm.10681 NP_056599.1	F:2.17 (YtoO)	NP_006466	osteoblast specific factor 2 (fasciclin I-like); periostin	1423	0
		AAN17733	extracellular matrix protein periostin-bm	1379	0
	·	S36111	osteoblast-specific factor 2	1372	0
		NP_000349	transforming growth factor, beta-induced, 68kDa; corneal dystroply, kerato-epithelin;	580	1E-165
			transforming growth factor, beta-induced, 68kD		
		AAC08449	BIGH3	497	1E-140
1					
A K 0 0 7 7 1 0 Mm.27385 BAB25204.1	F:2.17 (5to19)	NP_079012.1	079012.1 hypothetical protein FLJ12150	463	1E-129
		AAG22861.1	PKSG10	341	1E-95

Ŋ

7

1038 0		rotein 2; 1036 0	1036	1036	1036 968 966 441 IE-12	1036 968 966 441	1036 968 966 441 441	1036 968 966 441 441 439 393	1036 968 966 441 441 439 393	968 966 441 1 441 1 439 1 332	1036 968 966 441 1 441 1 439 1 322 323	1036 968 966 441 1 441 1 439 1 322 322	1036 968 966 441 1 441 1 439 1 322 322 323 728	1036 968 966 441 1 441 1 439 1 322 322 323 728	968 968 966 441 1 439 1 332 322 322 728	1036 968 966 441 1E-12 439 1E-12 393 1E-10 315 2E-8 715 715	1036 968 966 441 1E-12 441 1E-12 439 1E-10 393 1E-10 315 2E-8 315 2E-8 715 715 715
?) (DEAD-box, X isoform)		001347.2 DEAD/H (Asp-Glu-Ala-Asp/His) box polypeptide 3; DEAD/H box-3; helicase like protein 2;	; DEAD/H box-3; helicase like protei	DEAD/H (Asp-Glu-Ala-Asp/His) box polypeptide 3; DEAD/H box-3; helicase like protein 2; CAP-Rf DEAD/H (Asp-Glu-Ala-Asp/His) box polypeptide, Y chromosome; DEAD/H box-3, Y-linked DEAD-box protein 3, Y-chromosomal	; DEAD/H box-3; helicase like protein chromosome; DEAD/H box-3, Y-lin; VASA protein	; DEAD/H box-3; helicase like proteir 7 chromosome; DEAD/H box-3, Y-lin ; VASA protein	; DEAD/H box-3; helicase like protein ? chromosome; DEAD/H box-3, Y-lin ; VASA protein	DEAD/H (Asp-Glu-Ala-Asp/His) box polypeptide 3; DEAD/H box-3; helicase like protein 2; CAP-Rf DEAD/H (Asp-Glu-Ala-Asp/His) box polypeptide, Y chromosome; DEAD/H box-3, Y-linked DEAD/H (Asp-Glu-Ala-Asp/His) box polypeptide 4; VASA protein probable RNA helicase protein DKFZp434B1122.1 DEAD box RNA helicase similar to DEAD (aspartate-glutamate-alanine-aspartate) box polypeptide 3; D-E-A-D (aspartate-plusmate-alanine-aspartate) box polypeptide 3; D-E-A-D (aspartate-plusmate-alanine-aspartate)	; DEAD/H box-3; helicase like protein chromosome; DEAD/H box-3, Y-lin yASA protein te) box polypeptide 3; D-E-A-D (aspart ryonic RNA helicase 17 isoform 1; DEAD/H (Asp-Glu-,	2AP-Rf CAP-Rf CAP-Rf CAP-Rf DEAD/H (Asp-Glu-Ala-Asp/His) box polypeptide 3; DEAD/H box-3; helicase like protein 2; CAP-Rf DEAD/H (Asp-Glu-Ala-Asp/His) box polypeptide, Y chromosome; DEAD/H box-3, Y-linked DEAD-box protein 3, Y-chromosomal DEAD/H (Asp-Glu-Ala-Asp/His) box polypeptide 4; VASA protein probable RNA helicase protein DKFZp434B1122.1 probable RNA helicase protein DKFZp434B1122.1 DEAD/H (Asp-Glu-Ala-Asp/His) box polypeptide 3; embryonic RNA helicase glutamate-alanine-aspartate-glutamate-alanine-aspartate) box polypeptide 3; embryonic RNA helicase glutamate-alanine-aspartate) box polypeptide 17 isoform 1; DEAD/H (Asp-Glu-Ala-Asp/His) box polypeptide 17 (72kD); probable RNA-dependent helicase p72	; DEAD/H box-3; helicase like protein chromosome; DEAD/H box-3, Y-lin; VASA protein te) box polypeptide 3; D-E-A-D (aspartryonic RNA helicase 17 isoform 1; DEAD/H (Asp-Glu	; DEAD/H box-3; helicase like protein chromosome; DEAD/H box-3, Y-lin ; VASA protein te) box polypeptide 3; D-E-A-D (aspart ryonic RNA helicase 17 isoform 1; DEAD/H (Asp-Glu-4-dependent helicase p72	O01347.2 DEAD/H (Asp-Glu-Ala-Asp/His) box polypeptide 3; DEAD/H box-3; helicase like protein 2; CAP-Rf CAP-Rf O04651.2 DEAD/H (Asp-Glu-Ala-Asp/His) box polypeptide, Y chromosome; DEAD/H box-3, Y-linked DEAD-box protein 3, Y-chromosomal O61912.1 DEAD-H (Asp-Glu-Ala-Asp/His) box polypeptide 4; VASA protein probable RNA helicase protein DKFZp434B1122.1 F86585.1 DEAD box RNA helicase Blutamate-alanine-aspartate-glutamate-alanine-aspartate) box polypeptide 3; embryonic RNA helicase glutamate-alanine-aspartate) box polypeptide 3; embryonic RNA helicase D06377.1 DEAD/H (Asp-Glu-Ala-Asp/His) box polypeptide 17 isoform 1; DEAD/H (Asp-Glu-Ala-Asp/His) box polypeptide 17 (72kD); probable RNA-dependent helicase p72 Asp/His) box polypeptide 17 (72kD); probable RNA-dependent helicase p72 Asp/His) box polypeptide 18 protein 17	DEAD/H (Asp-Glu-Ala-Asp/His) box polypeptide 3; DEAD/H box-3; helicase like protein 2; CAP-Rf CAP-Rf 004651.2 DEAD/H (Asp-Glu-Ala-Asp/His) box polypeptide, Y chromosome; DEAD/H box-3, Y-linked 523 DEAD/H (Asp-Glu-Ala-Asp/His) box polypeptide 4; VASA protein 407 probable RNA helicase protein DKFZp434B1122.1 386585.1 DEAD box RNA helicase glutamate-alanine-aspartate-glutamate-alanine-aspartate) box polypeptide 3; DE-A-D (aspartate-glutamate-alanine-aspartate) box polypeptide 3; DE-AD/H (Asp-Glu-Ala-Asp/His) box polypeptide 17 (72kD); probable RNA-dependent helicase p72 Asp/His) box polypeptide 17 (72kD); probable RNA-dependent helicase p72 Asp/His) box polypeptide 17 (72kD); probable RNA-dependent helicase p72 Growth regulated nuclear 68 protein 17 O01068.1 UDP glycosyltransferase 2 family, polypeptide B17; UDP-glucuronyltransferase, family 2, beta-in 17 CHUG4) (HLUG4)	DEAD/H (Asp-Glu-Ala-Asp/His) box polypeptide 3; DEAD/H box-3; helicase like protein 2; CAP-Rf CAP-Rf O04651.2 DEAD/H (Asp-Glu-Ala-Asp/His) box polypeptide, Y chromosome; DEAD/H box-3, Y-linked DEAD/H (Asp-Glu-Ala-Asp/His) box polypeptide 4; VASA protein DEAD-box protein 3, Y-chromosomal O61912.1 DEAD/H (Asp-Glu-Ala-Asp/His) box polypeptide 4; VASA protein probable RNA helicase protein DKFZp434B1122.1 F86585.1 DEAD box RNA helicase protein DKFZp434B1122.1 Babb box RNA helicase protein DKFZp434B1122.1 Babb box RNA helicase O06377.1 DEAD/H (Asp-Glu-Ala-Asp/His) box polypeptide 3; embryonic RNA helicase Butamate-alauine-aspartate) box polypeptide 3; embryonic RNA helicase O06377.1 DEAD/H (Asp-Glu-Ala-Asp/His) box polypeptide 17 isoform 1; DEAD/H (Asp-Glu-Ala-Asp/His) box polypeptide 17 (72kD); probable RNA-dependent helicase p72 Asp/His) box polypeptide 17 (72kD); probable RNA-dependent helicase p72 O11097.5 growth regulated nuclear 68 protein O11097.5 similar to UDP-glucuronosyltransferase 2B15 precursor, microsomal (UDPGT) (UDPGT)H-3) (HLUG4) (HLUG4) (HLUG4) O11097.5 similar to UDP-glucuronosyltransferase 2 family, polypeptide B15; UDP-glucuronyltransferase, family 2, beta-Italy UDP glycosyltransferase 2 family, polypeptide B15; UDP-glucuronyltransferase, family 2, beta-Italy UDP glycosyltransferase 2 family, polypeptide B15; UDP-glucuronyltransferase, family 2, beta-Italy UDP glycosyltransferase 2 family, polypeptide B15; UDP-glucuronyltransferase, family 2, beta-Italy UDP glycosyltransferase 2 family, polypeptide B15; UDP-glucuronyltransferase, family 2, beta-Italy UDP glycosyltransferase 2 family, polypeptide B15; UDP-glucuronyltransferase, family 2, beta-Italy UDP glycosyltransferase 2 family, polypeptide B15; UDP-glucuronyltransferase, family 2, beta-Italy UDP glycosyltransferase 2 family, polypeptide B15; UDP-glucuronyltransferase, family 2, beta-Italy UDP glycosyltransferase 2 family, polypeptide B15; UDP-glucuronyltransferase, family 2, beta-Italy UDP-glycosyltransferase 2 family, polype	; DEAD/H box-3; helicase like protein / chromosome; DEAD/H box-3, Y-lin ; VASA protein iryonic RNA helicase 17 isoform 1; DEAD/H (Asp-Gludependent helicase p72 UDP-glucuronyltransferase, family 2, t UDP-glucuronyltransferase, family 2, t	O01347.2 DEAD/H (Asp-Glu-Ala-Asp/His) box polypeptide 3, DEAD/H box.3; helicase like protein 2, CAP-Rf O04651.2 DEAD/H (Asp-Glu-Ala-Asp/His) box polypeptide, Y chromosome; DEAD/H box.3, Y-linked DEAD/H (Asp-Glu-Ala-Asp/His) box polypeptide 4; VASA protein probable RNA helicase protein DK-FZp434B1122.1 DEAD/H (Asp-Glu-Ala-Asp/His) box polypeptide 4; VASA protein probable RNA helicase protein DK-FZp434B1122.1 BEAD/H (Asp-Glu-Ala-Asp/His) box polypeptide 3; embryonic RNA helicase DOG377.1 DEAD/H (Asp-Glu-Ala-Asp/His) box polypeptide 3; embryonic RNA helicase protein DEAD/H (Asp-Glu-Ala-Asp/His) box polypeptide 17 (72kD); probable RNA-dependent helicase p72 BINIMARIA (Asp-Glu-Ala-Asp/His) box polypeptide B17; UDP-glucuronyltransferase, family 2, beta-17 (11097.5) similar to UDP-glucuronosyltransferase 2 family, polypeptide B15; UDP-glucuronyltransferase, family 2, beta-15 DOG377.1 UDP glycosyltransferase 2 family, polypeptide B15; UDP-glucuronyltransferase, family 2, beta-15 S05093.1 UDP-glucuronosyltransferase 2B4 precursor, microsomal (UDPGT) (Hyodeoxycholic acid) (HLUG25) (UDP-glucuronosyltransferase 2B4 precursor, microsomal (UDPGT) (Hyodeoxycholic acid) (HLUG25) (UDPGTH-1)
DEAD-box protein 3 (Helicase-like protein 2) (HLP2) (DEAD-box, X isoform)		lypeptide 3; DEAD/H box-3; he	lypeptide 3; DEAD/H box-3; he	lypeptide 3; DEAD/H box-3; he lypeptide, Y chromosome; DEA	DEAD/H (Asp-Glu-Ala-Asp/His) box polypeptide 3; DEAD/H box-3; he CAP-Rf CAP-Rf DEAD/H (Asp-Glu-Ala-Asp/His) box polypeptide, Y chromosome; DEAD-box protein 3, Y-chromosomal DEAD/H (Asp-Glu-Ala-Asp/His) box polypeptide 4; VASA protein	lypeptide 3; DEAD/H box-3; he lypeptide, Y chromosome; DEA lypeptide 4; VASA protein 34B1122.1	lypeptide 3; DEAD/H box-3; he lypeptide, Y chromosome; DEA lypeptide 4; VASA protein 34B1122.1	lypeptide 3; DEAD/H box-3; he lypeptide, Y chromosome; DEA lypeptide 4; VASA protein 34B1122.1	DEAD/H (Asp-Glu-Ala-Asp/His) box polypeptide 3; DEAD/H box-3; he CAP-Rf CAP-Rf DEAD/H (Asp-Glu-Ala-Asp/His) box polypeptide, Y chromosome; DEA DEAD-box protein 3, Y-chromosomal DEAD-box protein 3, Y-chromosomal DEAD/H (Asp-Glu-Ala-Asp/His) box polypeptide 4; VASA protein probable RNA helicase protein DKFZp434B1122.1 DEAD box RNA helicase similar to DEAD (aspartate-glutamate-alanine-aspartate) box polypeptide 3; glutamate-alanine-aspartate) box polypeptide 3; glutamate-alanine-aspartate) box polypeptide 17 isoform 1; DEA DEAD/H (Asp-Glu-Ala-Asp/His) box polypeptide 17 isoform 1; DEA	DEAD/H (Asp-Glu-Ala-Asp/His) box polypeptide 3; DEAD/H box-3; helica CAP-Rf CAP-Rf DEAD/H (Asp-Glu-Ala-Asp/His) box polypeptide, Y chromosome; DEAD/F DEAD-box protein 3, Y-chromosonal DEAD-box protein 3, Y-chromosonal DEAD/H (Asp-Glu-Ala-Asp/His) box polypeptide 4; VASA protein probable RNA helicase protein DKFZp434B1122.1 DEAD box RNA helicase similar to DEAD (aspartate-glutamate-alamine-aspartate) box polypeptide 3; D-glutamate-alamine-aspartate) box polypeptide 3; EAD/A (Asp-Glu-Ala-Asp/His) box polypeptide 17 (72kD); probable RNA-dependent helicase p72	lypeptide 3; DEAD/H box-3; he lypeptide, Y chromosome; DEA lypeptide 4; VASA protein 34B1122.1 ine-aspartate) box polypeptide 3 olypeptide 17 isoform 1; DE. olypeptide 17 isoform 1; DE. obable RNA-dependent helicase	lypeptide 3; DEAD/H box-3; he lypeptide, Y chromosome; DEA lypeptide 4; VASA protein 34B1122.1 ine-aspartate) box polypeptide 3 ride 3; embryonic RNA helicase oolypeptide 17 isoform 1; DE.	lypeptide 3; DEAD/H box-3; he lypeptide, Y chromosome; DEA lypeptide 4; VASA protein 34B1122.1 anine-aspartate) box polypeptide 3 tide 3; embryonic RNA helicase oolypeptide 17 isoform 1; DE obable RNA-dependent helicase phide B17; UDP-glucuronyltrans	lypeptide 3; DEAD/H box-3; he lypeptide, Y chromosome; DEA lypeptide, Y chromosome; DEA lypeptide 4; VASA protein 34B1122.1 and aspartate) box polypeptide 3 tide 3; embryonic RNA helicase bolypeptide 17 isoform 1; DE bable RNA-dependent helicase ptide B17; UDP-glucuronyltrans ptide B17; UDP-glucuronyltrans 2B15 precursor, microsomal (U	lypeptide 3; DEAD/H box-3; he lypeptide, Y chromosome; DEA 34B1122.1 ine-aspartate) box polypeptide 3 tide 3; embryonic RNA helicase oolypeptide 17 isoform 1; DE obable RNA-dependent helicase piide B17; UDP-glucuronyltrans ptide B15; UDP-glucuronyltrans ptide B15; UDP-glucuronyltrans	lypeptide 3; DEAD/H box-3; he lypeptide, Y chromosome; DEA lypeptide 4; VASA protein 34B1122.1 une-aspartate) box polypeptide 3 tide 3; embryonic RNA helicase oolypeptide 17 isoform 1; DE bable RNA-dependent helicase ptide B17; UDP-glucuronyltrans ptide B15; UDP-glucuronyltrans ptide B15; UDP-glucuronyltrans	lypeptide 3; DEAD/H box-3; he lypeptide, Y chromosome; DEA anne-aspartate) box polypeptide 3 tide 3; embryonic RNA helicase olypeptide 17 isoform 1; DE shable RNA-dependent helicase ptide B17; UDP-glucuronyltransptide B15; UDP-glucuronyltransptid
Total In the property of Auto-Appel	sp/His) box polypeptide		an(Lie) how and amounted	sp/His) box polypeptide, hromosomal	sp/His) box polypeptide, hromosomal sp/His) box polypeptide	CAP-Rf DEAD/H (Asp-Glu-Ala-Asp/His) box polypeptide, DEAD-box protein 3, Y-chromosomal DEAD/H (Asp-Glu-Ala-Asp/His) box polypeptide 4 probable RNA helicase protein DKFZp434B1122.1	sp/His) box polypeptide, hromosomal sp/His) box polypeptide otein DKFZp434B1122.	sp/His) box polypeptide, hromosomal sp/His) box polypeptide otein DKFZp434B1122.	csp/His) box polypeptide, hromosomal csp/His) box polypeptide otein DKFZp434B1122glutamate-alanine-aspare) box polypeptide 3; en Asp/His) box polypeptide 3; en Asp/His) box polypeptide 3; en Asp/His) box polypeptide 3;	(sp/His) box polypeptide, hromosomal (sp/His) box polypeptide otein DKFZp434B1122. -glutamate-alanine-asparte) box polypeptide 3; en Asp/His) box polypeptide 3; en Asp/His) box polypeptide RN	sp/His) box polypeptide, hromosomal sp/His) box polypeptide otein DKFZp434B1122. sglutamate-alanine-aspar te) box polypeptide 3; en Asp/His) box polypeptide 3; en Asp/His) box polypeptide 68 protein	sp/His) box polypeptide, hromosomal sp/His) box polypeptide otein DKFZp434B1122glutamate-alanine-aspar te) box polypeptide 3; en Asp/His) box polypeptide 8; en Asp/His) box polypeptide 8; en 68 protein	csp/His) box polypeptide, hromosomal csp/His) box polypeptide otein DKFZp434B1122. e-glutamate-alanine-asparte) box polypeptide 3; en Asp/His) box polypeptide 3; en Asp/His) box polypeptide 8 protein 68 protein framily, polypeptide B17	sp/His) box polypeptide, hromosomal sp/His) box polypeptide otein DKFZp434B1122. e-glutamate-alanine-asparte) box polypeptide 3; en Asp/His) box polypeptide 3; en Asp/His) box polypeptide BN 17 (72kD); probable RN 68 protein family, polypeptide B17 family, polypeptide B17 syltransferase 2B15 prec	sp/His) box polypeptide, hromosomal sp/His) box polypeptide otein DKFZp434B1122glutamate-alanine-aspar te) box polypeptide 3; en Asp/His) box polypeptid 3; en Asp/His) box polypeptide BN 68 protein -family, polypeptide B17 -family, polypeptide B17 -family, polypeptide B15	sp/His) box polypeptide, hromosomal sp/His) box polypeptide otein DKFZp434B1122glutamate-alanine-aspar te) box polypeptide 3; en Asp/His) box polypeptide 3; en Asp/His) box polypeptide B17 (72kD); probable RN 68 protein -family, polypeptide B17 -family, polypeptide B15 -rase 2B15	sp/His) box polypeptide, hromosomal sp/His) box polypeptide otein DKFZp434B1122glutamate-alanine-aspar te) box polypeptide 3; en Asp/His) box polypeptide 3; en Asp/His) box polypeptide B17 (72kD); probable RN 68 protein -family, polypeptide B17 -family, polypeptide B15 -rase 2B15 -rase 2B15 -rase 2B15 -rase 2B15 -rase 2B15
box protein 3 (Helicas	H (Asp-Glu-Ala-Asp/l		f (u (A cm Ch, A la A cm)	CAP-Rf DEAD/H (Asp-Glu-Ala-Asp/His) box DEAD-box protein 3, Y-chromosomal	f H (Asp-Glu-Ala-Asp/l box protein 3, Y-chroi H (Asp-Glu-Ala-Asp/l	f H (Asp-Glu-Ala-Asp/l box protein 3, Y-chro H (Asp-Glu-Ala-Asp/l Ie RNA helicase protei	CAP-Rf DEAD/H (Asp-Glu-Ala-Asp/l DEAD-box protein 3, Y-chron DEAD/H (Asp-Glu-Ala-Asp/l probable RNA helicase protei DEAD box RNA helicase	f H (Asp-Glu-Ala-Asp/l -box protein 3, Y-chro H (Asp-Glu-Ala-Asp/l le RNA helicase protei box RNA helicase to DEAD (aspartate-glu	f H (Asp-Glu-Ala-Asp/I box protein 3, Y-chrof H (Asp-Glu-Ala-Asp/I le RNA helicase protei box RNA helicase to DEAD (aspartate-glu ate-alauine-aspartate) E H (Asp-Glu-Ala-Asp/	f H (Asp-Glu-Ala-Asp/l box protein 3, Y-chro H (Asp-Glu-Ala-Asp/l le RNA helicase box RNA helicase to DEAD (aspartate-glu ate-alauine-aspartate) E H (Asp-Glu-Ala-Asp/	CAP-Rf DEAD/H (Asp-Glu-Ala-Asp/His) bo DEAD-box protein 3, Y-chromoson DEAD/H (Asp-Glu-Ala-Asp/His) bo probable RNA helicase protein DKI DEAD box RNA helicase similar to DEAD (aspartate-glutamat glutamate-alanine-aspartate) box po DEAD/H (Asp-Glu-Ala-Asp/His) b Asp/His) box polypeptide 17 (72kD growtlı regulated nuclear 68 protein	f H (Asp-Glu-Ala-Asp/l -box protein 3, Y-chrof H (Asp-Glu-Ala-Asp/l Ie RNA helicase protei box RNA helicase to DEAD (aspartate-glu ate-alauine-aspartate) E H (Asp-Glu-Ala-Asp/ s) box polypeptide 17 regulated nuclear 68 p	f H (Asp-Glu-Ala-Asp/l box protein 3, Y-chron H (Asp-Glu-Ala-Asp/l le RNA helicase box RNA helicase to DEAD (aspartate-glu ate-alanine-aspartate) th H (Asp-Glu-Ala-Asp/ s) box polypeptide 17 regulated nuclear 68 p	f H (Asp-Glu-Ala-Asp/l box protein 3, Y-chron H (Asp-Glu-Ala-Asp/l le RNA helicase box RNA helicase to DEAD (aspartate-glu ate-alauine-aspartate) E H (Asp-Glu-Ala-Asp/ S) box polypeptide 17 regulated nuclear 68 p regulated nuclear 68 p to UDP-glucuronosylt to UDP-glucuronosylt	f H (Asp-Glu-Ala-Asp/I box protein 3, Y-chro H (Asp-Glu-Ala-Asp/I le RNA helicase protei box RNA helicase to DEAD (aspartate-glu ate-alanine-aspartate) E H (Asp-Glu-Ala-Asp/I s) box polypeptide 17 regulated nuclear 68 p regulated nuclear 68 p to UDP-glucuronosylt to UDP-glucuronosylt lycosyltransferase 2 fan lycosyltransferase 2 fan lycosyltransferase 2 fan	CAP-Rf DEAD/H (Asp-Glu-Ala-Asp/His) be DEAD-box protein 3, Y-chromoson DEAD-box Protein 3, Y-chromoson DEAD-H (Asp-Glu-Ala-Asp/His) be probable RNA helicase Similar to DEAD (aspartate-glutamate glutamate-alanine-aspartate) box pol DEAD/H (Asp-Glu-Ala-Asp/His) the Asp/His) box polypeptide 17 (72kD growtln regulated nuclear 68 protein UDP glycosyltransferase 2 family, por 17 Similar to UDP-glucuronosyltransfer (HLUG4) UDP glycosyltransferase 2 family, por 15 UDP-glucuronosyltransferase 2 family, por 15	CAP-Rf O04651.2 DEAD/H (Asp-Glu-Ala-Asp/His) box polype O61912.1 DEAD/H (Asp-Glu-Ala-Asp/His) box polype O61912.1 DEAD/H (Asp-Glu-Ala-Asp/His) box polype Applicate probable RNA helicase protein DKFZp434B E86585.1 DEAD box RNA helicase O66968.2 similar to DEAD (aspartate-glutamate-alanine-glutamate-alanine-aspartate) box polypeptide Asp/His) box polypeptide 17 (72kD); probab Asp/His) box polypeptide 17 (72kD); probab O01068.1 UDP glycosyltransferase 2 family, polypeptid (HLUG4) O01067.1 UDP glycosyltransferase 2 family, polypeptid 15 D55093.1 UDP-glucuronosyltransferase 2B15 acid) (HLUG25) (UDPGTH-1)
	1347.2 DEAD/H (551.2	551.2	651.2	912.1	651.2 912.1 85.1	651.2 912.1 85.1 968.2	651.2 912.1 85.1 968.2	651.2 912.1 85.1 968.2 377.1	651.2 912.1 85.1 968.2 377.1	CAP-Rf 4651.2 DEAD/h (CAP-Rf 4651.2 DEAD/h (CAP-Rf 4651.2 DEAD/h (CAP-Rf 4651.2 DEAD/H (CAP-Rf 4651.2 DEAD/H (3 DEAD-bo 1912.1 DEAD/H (7 probable F 585.1 DEAD bo 6968.2 similar to I glutamate- 6377.1 DEAD/H Asp/His) th 17 1068.1 UDP glycc 17 1097.5 similar to (HLUG4) 115 5093.1 UDP-glucc 13 6345.4 similar to I acid) (HL)
[00057]				NP_00465 015523		NP_00465 O15523 NP_06191 T46407	NP_004651.2 015523 NP_061912.1 T46407 AAF86585.1	NP_00465 015523 NP_06191 T46407 AAF8658 XP_06696	NP_00465 015523 NP_06191 T46407 AAF8658 XP_06696	NP_00465 015523 NP_06191 T46407 AAF8658 XP_06690	NP_00465 015523 NP_06191; T46407 AAF86585 XP_06696 NP_00637	NP_00465 015523 NP_06191 T46407 AAF8658 XP_06690 NP_00637	NP_00465 015523 NP_06191 T46407 AAF8658 XP_06696 NP_00637 NP_00637	NP_00465 015523 NP_06191 T46407 AAF8658 XP_06690 NP_00637 NP_00637 NP_00100 NP_01100	NP_00465 015523 NP_06191 T46407 AAF8658 XP_06690 NP_00637 NP_00100 XP_01100	NP_004651.2 015523 NP_061912.1 T46407 AAF86585.1 XP_066968.2 NP_006377.1 NP_001068.1 NP_011097.5 AAD55093.1	NP_00465 015523 NP_06191 T46407 AAF8658 XP_06696 NP_00637 NP_00100 NP_00100 NP_00100 NP_00100 NP_00100 XP_01100 XP_01303
F:2.18 (5to 19)													F:2.18 (5to19)	F:2.18 (5to19)	F:2.18 (5to19)	F:2.18 (5to19)	F:2.18 (5to19)
													Mm.160362 F:	Mm.160362 F:	Mm.160362 F:	Mm.160362 F:	Mm.160362 F:
NM_010028 Mm.88188	_ NP 034158.1												N M_053215 Mm.160362 F:2.18 (5to19)	N M_053215 Mi	VM_053215M	VM_053215 Mi	VM_053215 Miv 444445.1

Ŋ

0

<u>...</u>

 \sim

	•	0	0		0	0	0	0	0	0	0	1E-177	1E-168	1E-111		1E-110	1E-108	0	0
808		807	804		803	802	794	721	L89	655	635	620 1	590 1			397 1	390	 269	269
AP_000887.1 cytochrome P450, subfamily IVF, polypeptide 3: leukottieue B4 oueug hydroxylase- leukottiene-	B4 20-monooxygenase; cytochrone P450-LTB-oncga	Cytochrome P450 4F12 (CYPIVF12)	JP_009184.1 cytochrome P450, subfamily IVF, polypeptide 8; microsomal monooxygenase; flavoprotein-	linked monooxygenase	NP_076433.1 cytochrome P450 isoform 4F12	similar to cytochrome P450	hypothetical protein	F22329_1	(P_029070.2 similar to Cytochrome P450 4F12 (CYPIVF12)	BAC04868.1 unnamed protein product	XP_065069.2 similar to CYTOCHROME P450 4F6 (CYPIVF6)	XP_065068.1 similar to Cytochrome P450 4F12 (CYPIVF12)	cytochrone P-450LTBV	Cytochrome P450 4A11 precursor (CYPIVA11) (Fatty acid omega-hydroxylase) (P-450 HK	omega) (Lauric acid omega-hydroxylase) (CYP4AII) (P450-HL-omega)	overver i cycomoune rapo, suntaminy 1949, polypeptide 11; tatty acid omega-hydroxylase; P450HL-lonnega; alkane-1 monooxygenase; lauric acid omega-hydroxylase	fally acid omega-hydroxylase (EC 1.14.15) cytochrome P450 4A11 -	Similar to extracellular matrix protein 1	IP_004416.1 extracellular matrix protein 1, isoform 1 precursor; secretory component p85
NP_000887		Q9HCS2	NP_009184		NP_076433	AAH35350.1	CAD38795.1	AAC11543.1	XP_029070.	BAC04868.	XP_065069	XP_065068.	BAA02145.	Q02928	096000 dN	141 - 000 105.	165981	AAH23505.1	NP_004416.
																		F:2.18 (5to11)	
																		Mm.3433	·
																		NM_007899 Mm.3433 NP_031925.1	

C	V
C)
C	V

, 0,000,000 cm.		202	;	
NP_05/913.1				
	A34372	complement C6 precursor [validated]	1246	0
	XP_170508.	170508.1 similar to Complement component C6 precursor	916	0
	AAB59433.1	complement component C6	760	0
	NP_000578.	000578.1 complement component 7 precursor	397	1E-110
	CAA60121.1	complement C7	394	1E-109
NM_021525 Mm.28630 F:2.19 NP 067500.1	F:2.19 (7to19) AAH01025.1	AAH01025.1 Similar to RNA cyclase homolog	729	0
	Q9Y2P8	RNA 3'-terminal phosphate cyclase-like protein	727	0
	NP_005763.	.005763.2 RNA cyclase homolog	724	0
	AAF29016.1 HSPC338	HSPC338	637	0
	AAD32456.1	RNA cyclase homolog	514	1E-146
NM_022434 Mm.10976 F:2.19 NP_071879.1	F:2.19 (5to19) AAC08589.1	cytochrome P-450	855	0
	BAA75823.1	Leukotriene B4 omega-hydroxylase	855	0
	NP_001073.	001073.3 cytochrome P450, subfamily IVF, polypeptide 2; leukotriene B4 omega-hydroxylase; leukotriene- B4 20-	853	0
	Q9HBI6	Q9HBI6 *,	848	0
	NP_067010.	067010.1 cytochrome P450, subfamily IVF, polypeptide 11	848	0
	AAC50052.2	AACS0052.2 cytochrome P450 4F2	845	0
				_

			,	201		
				erythroblastic leukemia viral oncogene homolog 2 (neuro/glioblastoma derived oncogene		
				homolog)		
			AAH02706.1	Similar to v-erb-b2 avian erythroblastic leukemia viral oncogene homolog 3	292	2E-78
	-		AAD56009.2 herstatin	herstatin	283	9E-76
NM_021522 Mm.25149 NP_067497.1	25149	F:2.21 (5to19)	NP_005142.1	NP_005142.1 ubiquitin specific protease 14	910	0
NM_007711 Mm.28842		F:2.2 (YtoM)	AAB95161	chloride channel protein 3	1430	
NP 031737.1					7757	
			NP 001820	chloride channel 3; CIC-3	1415	
			P51790	Chloride channel protein 3 (CIC-3).	1403	0
				clcn3e	1368	0
			P51793	Chloride channel protein 4 (CIC-4).	1152	Te
	•		NP 001821.	chloride channel 4; CIC-4	1149	
				chloride channel 5; Chloride channel-5	1008	٦
				chloride channel protein, kichey - human	43.5	1117
					171	12-11/
NM 008245 Mm.33896		F:2.2 (7to19)	NP 002720 1	NP 002720 I hematonoietically expressed homeohow, proline rich homeof		
NP 032271.1		,	-	factor	381	1E-105
			JN0767	homeobox protein HEX	380	1E-105
			AAH14336.1	Similar to hematopoietically expressed homeobox	370	1E-105
				meobox related protein	246	2E-65
A K 0 0 3 1 2 1 Mm.30724	1	F:2.2 (5to19)	NP_076932.1	NP_076932.1 hypothetical protein MGC3279 similar to collectins	420	1E-116
BAB22581.1						
			NP_006429.1	collectin sub-family member 10; collectin liver 1; collectin 34	228	2E-58
	- 1					
NM 016704 Mm. 20247	1	F:2.2 (5to19)	NP 000056.1	P 000056.1 Complement component 6 precursor	1249	10

				NOON		
NM_007912 Mm.8534 F:2.21 (5to19) P00533	lm.8534	F:2.21 (5to19)		Epidernal growth factor receptor precursor (Receptor protein-tyrosine kinase ErbB-1)	1160	0
NP 031938.1						
			AAA52371.1	aberrant epidermal growth factor receptor	1160	0
			NP_005219.1	005219.1 epidermal growth factor receptor (erythroblastic leukemia viral (v-erb-b) oncogene homolog,	1157	0
_				avian); epidermal growth factor receptor (avian erythroblastic leukemia viral (v-erb-b) oncogene		
		·		homolog); Epidermal growth factor receptor		
			AAG35786.1	p110 epidermal growth factor receptor	1141	0
			AAG35790.1	AAG35790.1 truncated epidermal growth factor receptor	1141	0
			CAA25282.1	EGF (1 is 2nd base in codon)	942	0
			1007208A	epidermal growth factor receptor	884	0
			AAC50802.1	epidermal growth factor receptor precursor	700	0
			NP_005226.1	005226.1 v-erb-a erythroblastic leukemia viral oncogene homolog 4; avian erythroblastic leukemia viral (v-	626	IE-179
				erb-b2) oncogene homolog 4; v-erb-a avian crythroblastic leukemia viral oncogene homolog-like		
				4		
			NP_001973.1	001973.1 v-erb-b2 erythroblastic leukennia viral oncogene homolog 3 (avian); Transformation gene BRBB-	603	1E-172
				3; v-erb-b2 avian erythroblastic leukemia viral oncogene homolog 3		
				kinase-related transforming protein (erbB3) (EC 2.7.1) precursor	602	IE-172
			P21860	Receptor protein-tyrosine kinase erbB-3 precursor (c-erbB3) (Tyrosine kinase-type cell surface	602	1E-172
·				receptor HER3)		
			7	Chain A, Structure Of The Her3 (Erbb3) Extracellular Domain	602	1E-172
			P04626	Receptor protein-tyrosine kinase erbB-2 precursor (p185erbB2) (NEU proto-oncogene) (C-erbB-	569	1E-162
				2) (Tyrosine kinase-type cell surface receptor HER2) (MLN 19)		
		·	NP_004439.1	004439.1 v-erb-b2 erythroblastic leukemia viral oncogene homolog 2, neuro/glioblastoma derived oncogene	569	569 1E-162
				homolog; Avian erythroblastic leukemia viral (v-erb-b2) oncogene homolog 2; v-erb-b2 avian		

Ŋ

c	7	١	
C	7	١	
_	_	4	

NFD 061224 1	(1010) 1111	Title Love 122.1 Strattin	Strain	471	7-1
6.4.T. L				:	
		S38956	epithelial cell marker protein 1	468	1E-131
		AAH01550.1	Similar to stratifin	397	1E-110
		NP_003397.1	NP_003397.1 tyrosine 3/tryptophan 5 -monooxygenase activation protein, zela polypeptide; protein kinase C	342	6E-94
			inhibitor protein-1; phospholipase A2		
		NP_006817.1	NP_006817.1 (tyrosine 3/tryptophan 5 -monooxygenase activation protein, theta polypeptide; 14-3-3 protein tau	u 341	2E-93
		NP_003395.1	NP_003395.1 tyrosine 3-monooxygenase/tryptophan 5-monooxygenase activation protein, beta polypeptide; 14-	- 326	6E-89
			3-3 protein beta/alpha; Protein kinase Cinhibitor protein-1; Protein 1054		
		NP_036611.2	NP_036611.2 tyrosine 3-monooxygenase/tryptophan 5-monooxygenase activation protein, gamma polypeptide	e 319	7E-87
·		NP_003396.1	NP_003396.1 lyrosine 3/tryptophan 5 -monooxygenase activation protein, eta polypeptide	314	2E-85
		AAA35483.1 14-3-3n	14-3-3n	311	1E-84
		S38532	protein 14-3-3 eta chain	307	3E-83
	,				
NM_007825 Mm.4781 F NP_031851.1	F:2.22 (5to19)	AAC95426.1	AAC95426.1 oxysterol 7alpha-	640	0
		NP_004811.1	NP_004811.1 cytochrome P450, subfamily VIIB, polypeptide 1; oxysterol 7alpha-hydroxylase	640	0
		P22680	Cytochrome P450 7A1 (Cholesterol 7-alpha-monooxygenase) (CYPVII) (Cholesterol 7-alpha-hydroxylase)	<u> </u>	5E-85
NM_021354 Mm.41803 F	F.2.21 (7to19)	NP_001379.1	NP_001379.1 developmentally regulated GTP binding protein 2	653	0
		AAH00493.1	AAH00493.1 developmentally regulated GTP-binding protein 2	650	0
		AAH20803.1	developmentally regulated GTP binding protein 1	364	1E-100
		NP_004138.1	NP_004138.1 developmentally regulated GTP binding protein 1; neural precursor cell expressed,		
			developmentally down-regulated 3; developmentally regulated GTP-binding protein 1		

ហ

 \vdash

15

			198		
		AAL50342.1	brain-muscle-ARNT-like transcription factor 2d	545	1E-153
		AAF71306.1,	AAF71306.1, bHLH-PAS transcription factor MOP9	539	1E-152
		AAF71307.1	bHLH-PAS transcription factor MOP9	538	1E-151
		AAH00172.2	Similar to transcription factor BMAL2	536	1E-151
		BAB01485.1	transcription factor BMAL.2	533	1E-150
		JC5407	brain and muscle Ah receptor nuclear translocator-like protein, BMAL le	478	1E-133
		PC4288	brain and muscle Ah receptor nuclear translocator-like protein, BMAL1d	451	IE-125
		NP_001659.1	NP_001659.1 aryl hydrocarbon receptor nuclear translocator	328	3E-88
		CAD38953.1	hypothetical protein	323	1E-86
		Q9HBZ2	Aryl hydrocarbon receptor nuclear translocator 2 (ARNT protein 2)	318	3E-85
		AAH36099.1	Unknown (protein for MGC:33872)	318	3E-85
		NP_055677.1	NP_055677.1 aryl-hydrocarbon receptor nuclear translocator 2; KIAA0307 gene product, aryl hydrocarbon receptor nuclear translocator 2	317	8E-85
		AAC03365.1	aryl hydrocarbon receptor nuclear translocator; Arnt	235	2E-60
•					
NM_022985 Mm.1608 F:2. NP_075361.2	F:2.22 (7to19)	NP_061879.2	NP_061879.2 protein associated with PRK1; hypothetical protein; associated with PRK1	395	1E-110
		CAB66533.1	lıypothetical protein	389	1E-108
		NP_005998.1	zinc finger protein 216	239	2E-63
		XP_048461.1	similar to protein associated with PRK1; hypothetical protein; associated with PRK1	191	2E-60
	-	AAD17528.1 unknown	unknown	224	6E-59

S

		NP 000187	Indroxysteroid (11-hefa) dehydropenase 2	215	1E-55
		B48544	11 hera-hvdroxvaternid dehvdropenase 2	214	117.55
		T	9	í	
NM_008180 Mm.7504 F:2	F:2.23 (5to19)	NP_000169.1	_ 000169.1 glutathione synthetase	810	0
NM_007468 Mm.4533 F:2 NP 031494.1	F.2.22 (7to11)	P06727	Apolipoprotein A-IV precursor (Apo-AIV)	432	1E-120
		CAA31955.1	apolipoprotein A-IV	431	1E-119
	4	NP_000473.1	_000473.1 apolipoprotein A-IV precusor	428	1E-119
		LPHUA4	apolipoprotein A-IV precusor	426	1E-118
		AAA51748.1	apolipoprotein A-IV precursor	422	1E-117
		AAB59516.1	xB59516.1 apolipoprotein A-IV	277	2E-73
NM_007489 Mm.12177 F.2. NP 031515.1	F:2.22 (7to11)	NP_001169.2	_001169.2 aryl hydrocarbon receptor nuclear translocator-like	1219	0
		JC5405	brain and muscle Ah receptor nuclear translocator-like protein, BMAL1b	1218	0
		000327	BMAL 1 protein (Brain and muscle ARNT-like 1) (Member of PAS protein 3) (Basic-helix-loop helix-PAS orphan MOP3) (BHLH-PAS protein JAP3)	1130	0
		AAC51213.1	PAS protein 3	1102	0
		NP_064568.2	NP_064568.2 transcription factor BMAL2	260	1E-158
		AAL50339.1	brain-muscle-ARNT-like transcription factor 2a	557	1E-157
		AAL50340.1	brain-muscle-ARNT-like transcription factor 2b	551	1E-155
		AAL50341.1	brain-muscle-ARNT-like transcription factor 2c	545	1E-153

IJ

0

15

			196		
		AAA35835.1	FGF receptor-1 precursor	257	1E-67
		-	hasic fihroblast growth factor receptor protein.	257	1E-67
		NP_065680.1	065680.1 ret proto-oncogene isoform b; RET transforming sequence, hydroxyaryl-protein kinase; cadherin	253	2E-66
	/		family member 12, oncogene RET		
		CAA31408.1	ret tyrosine kinase (AA 1 - 860)	253	2E-66
		NP_066124.1	NP_066124.1 ret proto-oncogene isoforma; RET transforming sequence; hydroxyaryl-protein kinase; cadherin	253	2E-66
			family member 12; oncogene RET		
		NP_065681.1	065681.1 ret proto-oncogene isoform c; RET transforming sequence; hydroxyaryl-protein kinase; cadherin	253	2E-66
			family member 12; oncogene RET		
		AAH04257.1	AAH04257.1 ret proto-oucogene (multiple endocrine neoplasia MEN2A, MEN2B and medullary thyroid	252	3E-66
_		i	carcinoma 1, Hirschsprung disease)		
		AAA36786.1	tyrosine kinase	251	6E-66
		138153	gene retII protein - hunan	251	6E-66
		B34735	protein-tyrosine kinase (EC 2.7.1.112) (ret)	249	4E-65
		AAA36524.1	papillary thyroid carcinoma-encoded protein	249	4E-65
		JN0290	protein-tyrosine kinase (EC 2.7.1.112) (clone lambda-ret-1)	248	5E-65
		JN0291	protein-tyrosine kinase (EC 2.7.1.112) (clone lambda-ret-5)	248	5E-65
		CAB46483.1	RET tyrosine kinase receptor	248	6E-65
		AAA60266.1	RET tyrosine kinase/cAMP protein kinase A subunit RI	247	1E-64
		A39061	protein-tyrosine kinase (EC 2.7.1,112) FLT3 (fins homolog)	224	1E-57
		1FGKA	Chain A, Crystal Structure Of The Tyrosine Kinase Domain Of Fibroblast Growth Factor	223	2E-57
			Receptor 1	·	
		NP_075599.1	075599.1 [fibroblast growth factor receptor 1 isoform 9 precursor; fms-related tyrosine kinase-2; heparin-	216	2E-55
			binding growth factorreceptor; FMS-like tyrosine kinase 2; basic fibroblast growth factor receptor	,	
			1; N-sam tyrosine kinase; FLG protein; protein-tyrosine kinase; tyrosylprotein kinase;		
			hydroxyaryl-protein kinase		
NM_008290 Mni.8877	F:2.23 (YtoO)	NP_002144	lıydroxysteroid (17-beta) dehydrogenase 2	391	1E-109
NP 032316.1					
			11beta-hydroxysteroid dehydrogenase (EC 1.1.1.146) type 2	215	_1_
		AAH36780	hydroxysteroid (11-beta) dehydrogenase 2	215	8E-56

Ŋ

	-		195		
		AAA58470.1	growth factor receptor	270	2E-71
-		NP_000133.1	fibroblast growth factor receptor 3, isoform 1 precursor; protein-tyrosine kinase; tyrosylprotein	270	2E-71
			kinase; hydroxyaryl-protein kinase; human tyrosine kinase JTK4		
		AAM22078.1	fibroblast growth factor receptor 3	270	2E-71
		NP_075254.1	fibroblast growth factor receptor 3, isoform 2 precursor; protein-tyrosine kinase; tyrosylprotein	270	2E-71
			kinase; hydroxyaryl-protein kinase; human tyrosine kinase JTK4		-
		1GJOA	1GJOA Chain A, The Fgft2 Tyrosine Kinase Domain.	269	3E-71
		AAH15035.1	similar to fibroblast growth factor receptor 1 (fins-related tyrosine kinase 2, Pfeiffer syndrome)	258	5E-68
		AAA35837.1	libroblast growth factor receptor (FGFr) transmembrane form	257	1E-67
		NP_075594.1	075594.1 fibroblast growth factor receptor 1 isoform 4 precursor; fins-related tyrosine kinase-2; heparin-	257	1E-67
1			binding growth factorreceptor; FMS-like tyrosine kinase 2; basic fibroblast growth factor receptor		
			1; N-sam tyrosine kinase; FLG protein;protein-tyrosine kinase; tyrosylprotein kinase;		
			hydroxyaryl-protein kinase		
		NP_075593.1	fibroblast growth factor receptor 1 isoform 3 precursor; fins-related tyrosine kinase-2; heparin-	257	1E-67
			binding growth factorreceptor; FMS-like tyrosine kinase 2; basic fibroblast growth factor receptor		
	·	•	1; N-sam tyrosine kinase; FLG protein;protein-tyrosine kinase; tyrosylprotein kinase;		<u> </u>
			hydroxyaryl-protein kinase		
		CAA68679.1	A68679.1 Ityrosine kinase	257	1E-67
		CAA36101.1	436101.1 precursor polypeptide (AA -21 to 801)	257	1E-67
		NP_000595.1	000595.1 fibroblast growth factor receptor 1 isoform 1 precusor; fins-related tyrosine kinase-2; heparin-	257	1E-67
			binding growth factorreceptor; FMS-like tyrosine kinase 2; basic fibroblast growth factor receptor	•	
			1; N-sam tyrosine kinase; FLG protein;protein-tyrosine kinase; tyrosylprotein kinase;		
			hydroxyaryl-protein kinase		
		AAA35958.1	heparin-binding growth factor receptor	257	1E-67
		NP_056934.2	056934.2 fibroblast growth factor receptor 1 isoform 2 precursor; fins-related tyrosine kinase-2; heparin-	257	1E-67
			binding growth factorreceptor; FMS-like tyrosine kinase 2; basic fibroblast growth factor receptor		
			1; N-sam tyrosine kinase; FLG protein;protein-tyrosine kinase; tyrosylprotein kinase;		
			hydroxyaryl-protein kinase		
		AAH18128.1	418128.1 similar to fibroblast growth factor receptor 1 (fins-related tyrosine kinase 2, Pfeiffer syndrome)	257	1E-67

	SE-75			7E-75	7E-75	7E-75			9E-75	9E-75	9E-75	9E-75	1E-74	1E-74			1E-74	2E-74			2E-74	2E-74			7E-74	4E-72	4E-72			2E-71
	281			281	281	281			281	281	281	281	280	280			280	280			280	280			278	272	272			270
194	NP_000132.1 fibroblast growth factor receptor 2 isoform 1 precursor; keratinocyte growth factor receptor; K-	sam protein; proteintyrosine kinase, receptor like 14; FGF receptor; bacteria-expressed kinase;	fibroblast growth factor receptor BEK; tyrosylprotein kinase; hydroxyaryl-protein kinase	AAK94205.1 keratinocyte growth factor receptor 2 isoform BEK	AAK94209.1 keratinocyte growth factor receptor 2 isoform K-sam-IIC2	AAH39243.1 Similar to fibroblast growth factor receptor 2 (bacteria-expressed kinase, keratinocyte growth	factor receptor, craniofacial dysostosis 1, Crouzon syndrome, Pfeiffer syndrome, Jackson-Weiss	syndrome)	BAA89299,1 K-sam-IIO1	BAA89301.1 K-sam-IIO3	BAA89296.1 K-sam-IIH1	BAA89298.1 K-sam-IIH3	AAD31560.1 fibroblast growth receptor 2 IgIIIb isoform	NP_075262.1 fibroblast growth factor receptor 2 isoform 6 precursor; keratinocyte growth factor receptor; K-	sam protein; proteintyrosine kinase, receptor like 14; FGF receptor; bacteria-expressed kinase;	fibroblast growth factor receptor BEK; tyrosylprotein kinase; hydroxyaryl-protein kinase	A35969 heparin-binding growth factor receptor K-sam precursor	NP 075417.1 fibroblast growth factor receptor 2 isoform 10 precursor; keratinocyte growth factor receptor; K.	sam protein; proteintyrosine kinase, receptor like 14; FGF receptor; bacteria-expressed kinase;	fibroblast growth factor receptor BEK; tyrosylprotein kinase; hydroxyaryl-protein kinase	AAK94208.1 keratinocyte growth factor receptor 2 isoform K-sam-IIC3	NP_075261.1 fibroblast growth factor receptor 2 isoform 5 precursor; keratinocyte growth factor receptor; K-	sam protein; proteintyrosine kinase, receptor like 14; FGF receptor; bacteria-expressed kinase;	fibroblast growth factor receptor BEK; tyrosylprotein kinase; hydroxyaryl-protein kinase	AAD31561.1 fibroblast growth factor receptor 2 isoform IgIIIc isoform	BAC45037.1 isoform of FGFR2	NP_075259.1 fibroblast growth factor receptor 2 isoform 3 precusor; keratinocyte growth factor receptor; K-	sam protein; protein tyrosine kinase, receptor like 14; FGF receptor; bacteria-expressed kinase;	fibroblast growth factor receptor BEK; tyrosylprotein kinase; hydroxyaryl-protein kinase	AAM22079.1 fibroblast growth factor receptor 3

			193		
		NP 002244.1	kinase insert domain receptor (a type III receptor tyrosine kinase), Kinase insert domain receptor	326	2E-88
		NP_002011.1	NP_002011.1 fms-related tyrosine kinase 4, fms-related tyrosine kinase-4(vascular endothelial growth factor	325	3E-88
- :			receptor 3)		
		CAA43837.1	membrane protein	323	1E-87
		JC1402	protein-tyrosine kinase (EC 2.7.1.112) KDR - human	323	1E-87
		CAD27356.1	KIT protein	313	1E-84
		1VR2A	Chain A, Human Vascular Endothelial Growth Factor Receptor 2 (Kdr)Kinase Domain.	286	3E-76
		NP_075258.1	075238.1 fibroblast growth factor receptor 2 isoform 2 precursor; keratinocyte growth factor receptor; K-	284	8E-76
			sam protein; protein tyrosine kinase, receptor like 14; FGF receptor; bacteria-expressed kinase;		
			fibroblast growth factor receptor BEK; tyrosylprotein kinase; hydroxyaryl-protein kinase		
		NP_075264.2	075264.2 fibroblast growth factor receptor 2 isoform 8 precursor; keratinocyte growth factor receptor; K-	284	1E-75
			sam protein; protein tyrosine kinase, receptor like 14; FGF receptor; bacteria-expressed kinase;		
,			fibroblast growth factor receptor BEK; tyrosylprotein kinase; hydroxyaryl-protein kinase		
		NP_075419.1	075419.1 fibroblast growth factor receptor 2 isoform 12 precursor; keratinocyte growth factor receptor; K-	284	1E-75
			sam protein; proteintyrosine kinase, receptor like 14; FGF receptor; bacteria-expressed kinasc;		
			fibroblast growth factor receptor BEK; tyrosylprotein kinase; hydroxyaryl-protein kinase		
		A41794	keratinocyte growth factor receptor	284	1E-75
		AAK94206.1	keratinocyte growth factor receptor 2 isoform KGFR	284	1E-75
<u>-</u>	2003	NP_075420.1	fibroblast growth factor receptor 2 isoform 13 precursor; keratinocyte growth factor receptor; K-	283	1E-75
			sam protein; protein tyrosine kinase, receptor like 14; FGF receptor; bacteria-expressed kinase;		
			fibroblast growth factor receptor BEK; tyrosylprotein kinase; hydroxyaryl-protein kinase		
		BAA89300.1	K-sam-IIO2	283	2E-75
		NP_075263.1	075263.1 fibroblast growth factor receptor 2 isoform 7 precursor; keratinocyte growth factor receptor; K-	283	2E-75
	j		sam protein; proteintyrosine kinase, receptor like 14; FGF receptor; bacteria-expressed kinase;		
		ı	fibroblast growth factor receptor BEK; tyrosylprotein kinase; hydroxyaryl-protein kinase		
		Q01742	Fibroblast growth factor receptor BFR-2 precursor.	283	2E-75
		BAA89297.1	K-sam-11H2	281	5E-75
•		NP_075418.1	075418.1 fibroblast growth factor receptor 2 isoform 11 precursor; keratinocyte growth factor receptor; K-	187	SE-75
		-	sam protein; protein tyrosine kiņase, receptor like 14; FGF receptor; bacteria-expressed kinase;		
			fibroblast growth factor receptor BEK; tyrosylprotein kinase; hydroxyaryl-protein kinase		_

C	V
C	σ
r	-

NING 011656 Ndm, 10014	E:2 24 (11fe10)	MD 064512 1 Inflatin 1	192	681	0
	(21011) +7:7:1	141 - 004014:1		3	•
	,	AAH02933.1	Similar to tuftelin 1	622	1E-178
		BAB15615.1	unnamed protein product	271	3E-72
NM_021099 Mm.4394 NP 066922.1	F:2.23 (YtoO)	AAC50969.1	KIT protein	1532	0
		NP 000213.1	v-kit Hardy-Zuckerman 4 feline sarcoma viral oncogene homolog precursor	1526	0
		CAA27300.1	put. c-fins precursor	647	0
		NP_005202.2	NP_005202.2 colony stimulating factor 1 receptor precursor; FMS proto-oncogene; CD115 antigen; macrophage	647	0
			colony stimulating factor I receptor; similar to mouse Friend murine leukemia virus integration		
			site 2		
		AAH47521.1	colony stimulating factor 1 receptor, formerly McDonough feline sarcoma viral (v-fms) oncogene	645	0
			homolog		
		NP 006197.1	NP 006197.1 platelet-derived growth factor receptor alpha precursor	511	1E-144
		AAA36427.1	platelet-derived growth factor receptor	484	1E-136
	,	NP_002600.1	NP_002600.1 platelet-derived growth factor receptor beta precursor; beta platelet-derived growth factor	484	1E-136
		·	receptor		
		AAH32224.1	platelet-derived growth factor receptor, beta polypeptide	483	1E-135
	•	CAA81393.1	FLT3 receptor tyrosine kinase	442	1E-123
		NP 004110.1	fins-related tyrosine kinase 3	439	1E-122
		A36873	protein-tyrosine kinase (EC 2.7.1.112) STK-1 precursor	431	1E-120
		NP_002010.1	NP_002010.1 fins-related tyrosine kinase 1 (vascular endothelial growth factor/vascular permeability factor	358	5E-98
			receptor)		
		AAC16449.1	vascular endothelial growth factor receptor	357	7E-98
		P35916	Vascular endothelial growth factor receptor 3 precursor (VEGFR-3)(Tyrosine-protein kinase	327	8E-89
			receptor FLT4).		
		CAA48290.1	FTI_A	327	8E-89
		AAC16450.1	vascular endothelial growth factor receptor 2	326	2E-88

Ľ

 Ξ

12

 $\sqrt{2}$

				171		
						<u></u>
NM_013585 Mm.16251	Mm. 16251	F:2.25 (5to19)	S27332	proteasome endopeptidase complex (EC 3.4.25.1)	272	5E-73
NP 038613.1						
			NP_002791.1	NP_002791.1 proteasome beta 9 subunit isoform 1 proprotein; proteasome subunit, beta type, 9; proteasome-	269	4E-72
				related gene 2; proteasome chain 7; macropain chain 7; low molecular mass protein 2;		
		-		multicatalytic endopeptidase complex chain 7; proteasome catalytic subunit 1i; proteasome		
		·		subunit beta 6i		
	•		NP_683756.1	NP_683756.1 proteasome beta 9 subunit isoform 2 proprotein; proteasome subunit, beta type, 9; proteasome-	256	3E-68
				related gene 2; proteasome chain 7; macropain chain 7; low molecular mass protein 2;		
				multicatalytic endopeptidase complex chain 7; proteasome catalytic subunit 1i; proteasome		
				subunit beta 6i		
		-				
NM_008035 Mm.2724 NP 032061.1	Mm.2724	F:2.24 (YtoO)	CAA49267.1	CAA49267.1 folate receptor	421	1E-116
			NP_000794.1	NP_000794.1 folate receptor 2 precursor	419	1E-116
			AAA17370.1	AAA17370.1 folate binding protein	419	1E-116
			NP_000795.1	folate receptor 3 precursor	400	1E-110
			NP_000793.1	NP_000793.1 folate receptor 1 (adult)	374	1E-102
			AAA74896.1	AAA74896.1 folate-binding protein	368	1E-100
			XP_169247.1	XP_169247.1 similar to Folate receptor gamma precursor (FR-gamma) (Folate receptor 3)	313	4E-84
			AAB81937.1	folate binding protein	164	4E-74
NM_025649 Mm.27787	Mm.27787	F:2.24 (YtoO)	NP_055443	gene predicted from cDNA with a complete coding sequence; caught by MAD Two 2	424	1E-117
NP 079925.1						

Ŋ

 \Box

., ...

			190		
		NP_059112.1	059112.1 interleukin 1 receptor accessory protein-like 2; interleukin 1 receptor 9; IL-1 receptor; X-linked	506	3E-51
			interleukin-1 receptor accessory protein-like 2; IL-1 receptor accessory protein-like 2	•	
		AAF59412.1	X-linked interleukin-1 receptor accessory protein-like 2	206	3E-51
NM_008295 Mm.17910 F	F:2.25 (YtoO)	NP_000853	hydroxy-delta-5-steroid dehydrogenase, 3 beta- and steroid delta-isomerase 1; Hydroxy-delta-5-	528	1E-149
NP 032321.1			steroid dehydrogenase, 3 beta- and steroid		
		AAA51831	3-beta-hydroxysteroid dehydrogenase/delta-5-delta-4-isomerase.	526	1E-149
		NP_000189	hydroxy-delta-5-steroid dehydrogenase, 3 beta- and steroid delta-isomerase 2, Hydroxy-delta-5-	513	1E-145
			steroid dehydrogenase, 3 beta- and steroid		
		AAA36001	3-beta-hydroxysteroid dehydrogenase gene	481	1E-136
		CAC19801	dJ871G17.4 (novel 3-beta hydroxysteroid dehydrogenase/isomerase family member)	360	3E-99
		AAM08704.	3-beta-hydroxysteroid dehydrogenase	353	5E-97
	ال والمحاول الذي المحاول المحا	XP 060827	similar to 3-beta-hydroxysteroid dehydrogenase	258	2E-68
		AAG37824	3 beta-hydroxy-delta 5-C27-steroid oxidoreductase	225	2E-58
		NP 079469	3 beta-hydroxy-delta 5-C27-steroid oxidoreductase	223	8E-58
		CAC19803	dJ871G17.6 (novel 3-beta hydroxysteroid dehydrogenase/isomerase family member)	202	8E-52
		AAD14414	3 beta-hydroxysteroid dehydrogenase homolog pseudogene	199	7E-51
NM_008340 Mm.3135 F	F:2.25 (7to19)	NP_004961.1	004961.1 insulin-like growth factor binding protein, acid labile subunit; INSULIN-LIKE GROWTH	805	0
NP 032366.1			FACTOR BINDING PROTEIN COMPLEX ACID LABILE CHAIN PRECURSOR		
		AAH25681.1	insulin-like growth factor binding protein, acid labile subunit	804	0
NM_008343 Mm.29254 F NP 032369.1	F:2.25 (5to11)	NP_000589.1	_000589.1 insulin-like growth factor binding protein 3	442	1E-123
		NP_000590.1	insulin-like growth factor binding protein 5	506	4E-52
29 Mm.12616	F:2.25 (5to 19)	JC4775	interferon-induced double-stranded RNA-activated protein kinase inhibitor	897	0
NP 032955.1					
		AAH33823.1	Similar to DnaJ (Hsp40) homolog, subfamily C, member 3	213	8E-55

Ŋ

			189		
NM_028740 Mm.159128 F:2.27 (5to19)	F:2.27 (5to 19)	AAA51560.1	AAA51560.1 alpha-1-antichymotrypsin precursor	206	4E-53
NP 083016.1					
			alpha1-antichymotrypsin	206	4E-53
		pdb 1QMN	Chain A, Alphal-Antichymotrypsin Serpin In The Delta Conformation (Partial Loop Insertion)	206	4E-53
		AAH34554.1	serine (or cysteine) proteinase inhibitor, clade A (alpha-1 antiproteinase, antitrypsin), member 3	206	4E-53
		XP_028322.1	XP_028322.1 similar to Alpha-1-antichymotrypsin precursor (ACT)	206	4E-53
		CAA25459.1	alpha 1 antichymotrypsin	206	4E-53
		AAD08810.1	alpha-1-antichymotrypsin precursor	206	4E-53
		NP_001076.1	001076.1 alpha-1-autichymotrypsin, precursor; alpha-1-autichymotrypsin; antichymotrypsin	202	6E-52
		гтнис	alpha-1-antichymotrypsin precursor	202	6E-52
		1313184C	chymotrypsin inhibitor	202	6E-52
,					
75 Mm.29286	F:2.26 (YtoO)	NP_057066	germ cell specific Y-box binding protein; contrin	285	1E-76
NP 058571.1	!				
		AAH33800	germ cell specific Y-box binding protein	285	1E-76
			•		
NM_008362 Mm.896	F:2.26 (5to 19)	NP_000868.1	_000868.1 interleukin 1 receptor, type I precursor; interleukin 1 receptor alpha, type I; interleukin receptor	r 823	0
NP 032388.1			1; antigen CD121a		
		pdb 11RA	Interleukin-1 Receptor With The Interleukin-1 Receptor Antagonist (IIIRa)	451	1E-125
		pdb/11TB	Typc-1 Interleukin-1 Receptor Complexed With Interleukin-1 Beta	448	1E-124
		pdb 1G0Y	II-1 Receptor Type 1 Complexed With Antagonist Peptide Af10847	445	1E-123
		XP_002685.3		356	2E-96
		AAG21368.1	IL-1Rrp2	356	2E-96
		NP_003845.1	. 003845.1 interleukin 1 receptor-like 2	354	7E-96
		NP_057316.2	057316.2 interleukin 1 receptor-like 1; interleukin 1 receptor 1; ST2V protein	208	5E-52
	-				

		O9P255	Hypothetical zinc fineer protein KIAA1473	313	0E-85
		42.1	BAB85542.1 KIAA1956 protein	313	9E-85
		XP_085836.1	XP_085836.1 similar to Hypothetical zinc finger protein KIAA1956	313	9E-85
		XP_047550.1	XP_047550.1 similar to Hypothetical zinc finger protein KIAA1473	313	9E-85
		NP_003406.1	NP_003406.1 zinc finger protein 268	312	1E-84
		AAH36038.1	AAH36038.1 Unknown (protein for MGC:33240)	312	1E-84
		AAK69307.1 ZNF268B	ZNF268B	312	1E-84
		S47071	finger protein HZF3, Krueppel-related - human (fragment)	312	2E-84
		NP_037512.1	NP_037512.1 zinc finger protein 228	312	2E-84
NM_007517 Mm.2146	F:2.27 (7to19)	AAD43017.1	AAD43017.1 ancient ubiquitous 46 kDa protein AUP1	549	1E-155
NP 031543.1					
•	·	AAD43010.1	AUP1 homolog	548	1E-155
		BAB14753.1	unnamed protein product	492	1E-139
		NP_036235.1	NP_036235.1 ancient ubiquitous protein 1	431	1E-119
				\dagger	
NM_018816 Mm.2161 NP 061286.1	F:2.27 (5to19)	XP_165719.1	XP_165719.1 similar to Apolipoprotein M (ApoM) (G3a) (HSPC336)	319	3E-87
		CAB51604.1	G3a protein	299	2E-81
		AAF29014.1 HSPC336	HSPC336	262	4E-70

	5E-86	5E-86	6E-86	6E-86	6E-86	6E-86	6E-86	6E-86	8E-86	8E-86	1E-85	1E-85	.2E-85	2E-85	2E-85	2E-85	3E-85	3E-85	4E-85	5E-85
	317	317	317	317	317	317	317	317	316	316	315	315	315	315	315	315	314	314	314	313
187	NP_003442.1 zinc finger protein 177	XP_087503.1 similar to zinc finger protein 91 (HPF7, HTF10)	XP_033888.3 similar to Zinc finger protein 41	CAC88162.1 bB479F17.3 (zinc finger protein 41)	NP_700359.1 zinc finger protein 41	A54661 zinc finger protein ZNF41 - human	AAI-122992.1 Unknown (protein for MGC:29879)	XP_166367.1 similar to Zinc finger protein 184	BAC04216.1 unnamed protein product	NP_065704.1 zinc finger protein 287	NP_061025.3 zinc finger protein 331; zinc finger protein 463; C2H2-like zinc finger protein	AAF78075.1 KRAB zinc finger protein	AAH36714.1 Unknown (protein for IMAGE:4846514)	T12489 hypothetical protein DKFZp572P0920.1 - human	XP_032810.1 similar to Zinc finger protein 20 (Zinc finger protein KOX13) (DKFZp572P0920)	AAF88107.1 Hypothetical zinc finger-like protein	NP_612203.1 TRAF6-inhibitory zinc finger protein; TRAF6-binding zinc finger protein	XP_092088.3 similar to zinc finger protein 91 (FIPF7, HTF10)	XP_047554.4 sinular to Hypothetical zinc finger protein KIAA1473	NP_006620.1 zinc finger protein 271

	X	XP_092090.2	092090.2 similar to Hypothetical zinc finger protein KIAA1473	320	6E-87
·	B.	BAA24050.1	Zinc-finger protein	320	6E-87
	Z	P_443092.1	NP_443092.1 kruppel-like zinc finger protein	319	1E-86
	X	XP_171940.1	171940.1 similar to BC37295_1	318	2E-86
	Z	NP_653294.1	653294.1 hypothetical protein FLJ30932	318	2E-86
	X	XP_064929.5	064929.5 similar to Zinc finger protein 20 (Zinc finger protein KOX13)	318	2E-86
	Z	P_079009.1	079009.1 hypothetical protein FLJ14345	318	2E-86
	Z	NP_003416.1	003416.1 zinc finger protein 45 (a Kruppel-associated box (KRAB) domain polypeptide); Zinc finger protein-45 (a Kruppel-associated box (KRAB) domain	318	2E-86
	Ā	F63030.1	Zinc finger protein ZNF45	318	2E-86
	Z	NP_008889.1	008889.1 zinc finger protein 16 (KOX 9)	318	2E-86
	J.	CAD39111.1	hypothetical protein	318	2E-86
	X	XP_092093.1	092093.1 similar to Zinc finger protein 85 (Zinc finger protein HPF4) (HTF1)	318	2E-86
	Ы	17020	Zinc finger protein 16 (Zinc finger protein KOX9)	318	2E-86
-	Y	AH06528.1	AAH06528.1 zinc finger protein 43 (HTF6)	318	3E-86
	Х	XP_086128.1	086128.1 similar to Zinc finger protein 35 (Zfp-35)	318	3E-86
	X	XP_065116.3 s	similar to zinc finger protein 91 (HPF7, HTF10)	317	4E-86
	Z	NP_003414.1	.003414.1 zinc finger protein 43 (HTF6) [317	4E-86
	A	AAH35579.1	Similar to zinc finger protein 208	317	4E-86
	Z	P_061121.1	NP_061121.1 zinc finger protein ZFP	317	SE-86
				1	

		185		
	XP_091983.1	991983.1 similar to Zinc finger protein 135	324	4E-88
	CAB94232.2	zinc finger protein	323	5E-88
	NP_003419.1	003419.1 zinc finger protein 84 (HPF2)	323	5E-88
	B32891	finger protein 2, placental	323	5E-88
	NP_055295.1	055295.1 zinc finger protein AF020591	323	7E-88
,	AAC51180.1	kruppel-related zinc finger protein	323	9E-88
	XP_092097.1	092097.1 similar to Zinc finger protein 93 (Zinc finger protein HTF34)	322	1E-87
	AAH36110.1	AAH36110.1 Similar to zinc finger protein 208	322	1E-87
	BAC04610.1	unnamed protein product	322	2E-87
	NP_612143.1	612143.1 hypothetical protein FLJ31526	322	2E-87
	NP_067039.1	zinc finger protein 71; endothelial zinc finger protein induced by tumor necrosis factor alpha	321	3E-87
		003399.1 zinc finger protein 37 homolog (mouse); Zinc finger protein-37, mouse, homolog of; zinc finger protein homologous to Zfp37 in mouse	321	3E-87
	BAC04064.1	unnamed protein product	321	3E-87
	Q9Y6Q3	Zinc finger protein ZFP-37	321	3E-87
	AAD23608.1	AAD23608.1 BC37295_2 (partial)	321	3E-87
	AAL58442.1	zinc finger protein 328	321	3E-87
	BAB47481.1	KIAA1852 protein	321	3E-87
	AAH37209.1	Unknown (protein for MGC:41936)	320	4E-87
	XP_171752.1	171752.1 similar to zinc finger protein 29	320	4E-87

	1.84		
XP_031283.1	XP_031283.1 similar to Hypothetical zinc finger protein KIAA1710	337	3E-92
P35789	ein HTF34)	336	8E-92
NP_003427.1	NP_003427.1 zinc finger protein 135 (clone pHZ-17)	336	1E-91
NP_653290.2	653290.2 hypothetical protein FLJ32191	335	1E-91
BAB71257.1	unnamed protein product	333	8E-91
BAC04764.1	umamed protein product	332	2E-90
NP_003420.1	_003420.1 ziuc finger protein 85 (HPF4, HTF1)	330	5E-90
BAA86512.1	BAA86512.1 KJAA1198 protein	328	3E-89
XP_032674.1	032674.1 similar to Hypothetical zinc finger protein KIAA1198	328	3E-89
NP_660338.1	660338.1 similar to Zinc finger protein 136	327	4E-89
BAB71272.1	BAB71272.1 uunamed protein product	327	6E-89
XP_065387.2	XP_065387.2 similar to Zinc finger protein 135	326	1E-88
XP_086070.1	086070.1 similar to Zinc finger protein 93 (Zinc finger protein HTF34)	325	1E-88
NP_003421.1	NP_003421.1 zinc finger protein 91 (HPF7, HTF10)	325	1E-88
XP_068538.2	_068538.2 similar to Zinc finger protein 93 (Zinc finger protein HTF34)	325	2E-88
XP_028314.1	XP_028314.1 similar to KRAB zinc finger protein KR18	324	3E-88
XP_115658.2	XP_115658,2 similar to Zinc finger protein 208	324	3E-88
T14757	hypothetical protein DKFZp572C163.1	324	3E-88

NP_003992.2 Fc fragment of IgG, low affinity IIb, representer for (CD32)	Fe fragment of IgG, low affinity IIb, re	P_003992.2 Fc fragment of 1gG, low affinity IIb, receptor for (CD32); Fc fragment of 1gG, low affinity II,	296	4E-80
	JL0119	receptor tot (CD32) Fe gamma (IgG) receptor IIb precursor	296	4E-80
NM_029813 Mm.159813 F:2.28 (5to19) NP 084089.1	NP_689814.1	NP_689814.1 hypothetical protein FLJ38281	373	1E-103
	XP_091960.1	XP_091960.1 similar to zinc finger protein 14 (KOX 6); GIOT-4 for gonadotropin inducible transcription repressor-4	373	1E-103
	NP_066358.1	NP_066358.1 zinc finger protein 14 (KOX 6); GIOT-4 for gonadotropin inducible transcription repressor-4	364	1E-100
	NP_699189.1	NP_699189.1 hypothetical protein FLJ90396	364	1E-100
	XP_091958.1	XP_091958.1 similar to zinc finger protein 14 (KOX 6); GIOT-4 for gonadotropin inducible transcription repressor-4	364	1E-100
	XP_091968.4	XP_091968.4 similar to zinc finger protein 91 (HPF7, HTF10)	353	6E-97
	AAF71790.1	ZNF180	347	6E-95
	NP_037388.1	NP_037388.1 zinc finger protein 180 (HHZ168)	347	6E-95
	NP_003428.1	NP_003428.1 zinc finger protein 136 (clone pHZ-20)	345	1E-94
	NP_689815.1	NP_689815.1 hypothetical protein FLJ40981	344	3E-94
	NP_085116.1	NP_085116.1 hypothetical protein FLJ21628	343	8E-94
	AAD23607.1	BC37295_1	341	2E-93
	BAC04309.1		338	2E-92
	BAB21801.1	KIAA1710 protein	337	3E-92
	XP_032812.1	XP_032812.1 similar to hypothetical protein FLJ40981	337	3E-92

			182		
		P07864	L-lactate dehydrogenase C chain (LDH-C) (LDH testis subunit) (LDH-X)	494	1E-139
		pdb 110Z	Chain A, Human Heart L-Lactate Dehydrogenase H Chain, Ternary Complex With Nadh And	493	1E-139
			Oxamate		
		NP_149972.1	NP_149972.1 lactate dehydrogenase A -like	490	1E-138
		BAB71710.1	BAB71710.1 unnamed protein product	488	1E-137
		AAA59508.1	lactate deliydrogenase-C	437	1E-122
		XP_062669.6	XP_062669.6 similar to lactate dehydrogenase A -like	386	1E-107
		NP_659409.1	NP_659409.1 hypothetical protein MGC23940	343	3E-94
NM_010187 Mm.10809 F:2 NP_034317.1	F:2.28 (7to19)	AAD00638.1	Fc-gamma-RIIb2	309	6E-84
		AAD00641.1	Fc-gamna-RI1b2	306	6E-63
		CAA36713.1	precursor polypeptide (AA -42 to 249)	305	8E-83
		AAA35842.1	IgG Fe fragment receptor precursor	304	2E-82
		AAA36051.1	IgG Fc receptor beta-Fc-gamma-RII	301	1E-81
		CAA35644.1	IgG Fc receptor	301	2E-81
		AAD00639.1	Fc-gamma-RIIb1	301	2E-81
		AAD00637.1	Fc-gamma-RUb1	301	2E-81
		AAD00640.1	Fc-gannna-RIIb 1	297	2E-80
		P31994	Low affinity immunoglobulin gamma FC region receptor II-B precursor (FC-gamma RUI-B) (FCRII-B) (IGG FC receptor II-B) (FC-gamma-RIIB) (CD32) (CDW32)	297	3E-80

	0.	0	0	0	0	1E-63	1E-138	1E-138	4E-89	4E-88	1E-81	1E-170	1E-170	1E-140	1E-140	1E-139
	720	664	099	655	645	243	489	488	525	322	301	597	595	495	495	494
181	NP_004963.1 Janus kinase 2; tyrosine-protein kinase JAK2	JAK3_HUMAN; JANUS KINASE 3; JAK-3; LEUKOCYTE JANUS KINASE; L-JAK	JAK3	NP_000206.1 Janus kinase 3; L-JAK	JAK3B	Similar to Janus kinase 3 (a protein tyrosine kinase, leukocyte)	NP_005527.1 inositol(myo)-1(or 4)-monophosphatase 1	Inositol-1(or4)-monophosphatase (IMPase) (IMP) (Inositol monophosphatase) (Lithium-sensitive myo-inositol monophosphatase A1)	NP_055029.1 inositol(myo)-1(or 4)-monophosphatase 2	XP_095533.1 similar to Myo-inositol-1(or 4)-monophosphatase (IMPase) (IMP) (Inositol monophosphatase) (Lithium-sensitive myo-inositol monophosphatase A1)	brain myo-inositol monophosphatase A2b; IMPase A2b	NP_005557.1 lactate dehydrogenase A	Chain A, Human Muscle L-Lactate Dehydrogenase M Chain, Ternary Complex With Nadh And Oxamate	NP_002291.1 lactate dehydrogenase B	lactate dehydrogenase (E.C. 1.1.1.27)	NP_002292.1 lactate dehydrogenase C
	NP_004963.1	AAD22741.1	AAC50950.1	NP_000206.1	AAC50226.1	AAH28068.1	NP_005527.	рдр 2ННМ	NP_055029,1	XP_095533.1	AAF07824.1	NP_005557.	pdb 1110	NP_002291.	AAAS9507.1	NP_002292.
							NM_018864 Mm.183042 F.2.29 (5to19) NP 061352.1					NM_010699 Mm.141443 F:2.28 (7to19)	NP 034829.1	,		

Ŋ

7(

--1

 $\ddot{\circ}$

XP_035037.2 similar to MEGF7	simila	r to MEGF7	200	2E-50
A K 0 0 5 0 4 9 Mm.158752 F:2.3 (5to19)		AAH31569.1 Sinuilar to RIKEN cDNA 1300018K11 gene	899	0
	P22792	Carboxypeptidase N 83 kDa chain (Carboxypeptidase N regulatory subunit)	490	1E-137
C O C C C C C C C C C C C C C C C				
BAB26559.1		NF_U00808.1 endopiasuuc retteutum protein 29 precursor, endopiasmic retteutum lumenai protein EKp28	430	1E-119
NM_009547 Mm.29434 F:2.29 (5to7)		NP_003400.2 zinc finger protein 161 homolog; zinc finger protein homologous to Zfp161 in mouse	839	0
				·
	043829	Zinc finger protein Zfp-161 (Zinc finger protein 5) (hZF5)	838	0
	XP_008796.5	XP_008796.5 sinular to zinc finger protein 161 homolog (mouse); zinc finger protein homologous to Zfp161	288	2E-77
		in mouse		
NM_018793 Mm.20249 F:2.29 (5to19)		XP_008893.4 similar to Non-receptor tyrosine-protein kinase TYK2	1854	0
	AAH14243.1	AAH14243.1 Unknown (protein for MGC:20776)	1853	0
	NP_003322.1	NP_003322.1 tyrosine kinase 2	1839	0
	AAB22747.1	AAB22747.1 IFN-tyk, tyk2=interferon alpha/beta signaling pathway-related protein tyrosine kinase [human,	, 1442	0
		Daudi cell line, Peptide Partial, 899 aa)		
	A39577	A39577 protein-tyrosine kinase (EC 2.7.1.112) JAK1	924	0
	NP_002218.1	janus kinase 1	924	0
	AAC23653.1 Jak2 kinase	Jak2 kinase	720	0
	JW0091	Janus kinase (EC 2.7) 2 -	720	0

Ŋ

 \mathcal{L}

(1)

NP_0049183.1 Similar to Interferon-induced protein with terraticopeptide repeats 4 (IEIT.4) (Interferon-induced 509 1E-142	
204 504 504 500 365 333 2885 279 279 2161 2161 787 787 787	prince com
204 1 504 1 504 1 504 1 504 1 365 1 2285 22165 21140 21140 788 788 785	
204 1 504 1 500 1 365 1 333 285 285 2165 2161 2140 787 785	XP_048183.1 similar to
504 1 500 1 365 1 333 285 284 284 279 2165 2161 2161 2140 2140 788 787 785 785	_
285 285 284 284 279 279 2161 2161 71138 787 787 787	H04977.1
365 1 285 285 285 284 279 2161 2140 1138 787 785	NP_001540.1 interfere
ptide repeats 2 333 ats 1; Interferon, alpha-inducible protein 284 279 279 2165 2167 313 313 313 313 313 313 313 313 314 317 318 319 319 319 319 319 319 319 319 319 319	XP_084477.1 similar
ptide repeats 2 333 ats 1; Interferon, alpha-inducible protein 284 279 279 2165 2161 2167 313 313 314 317 318 319 319 319 319 319 319 319 319 319 319	
ats 1; Interferon, alpha-inducible protein 284 279 279 2165 2161 2140 2140 788 n) 785	H32839.1
ats 1; Interferon, alpha-inducible protein 284 279 279 2165 2161 2161 2178 n) 788	AAH07091.1 Unknov
279 279 2165 2161 2140 1138 n) 788	NP_001539.1 interfer
279 2165 2167 2161 2161 2170 1138 1138 1138 1138 1138 1138	(MW S
2165 2161 2140 1138 1138 788 787 787 771	NP_036552.1, retinoic
2165 2161 2140 1138 788 787 787 771	
2161 2140 1138 788 787 787 787 787	MMHUND nidoger
2 (osteonidogen); nidogen 2 2 (osteonidogen); nidogen 2 1-2 precursor (NID-2) (Osteonidogen). 787 787 787 o Nidogen-2 precursor (NID-2) (Osteonidogen) 787	NP_002499,1 nidoger
2 (osteonidogen); nidogen 2 788 -2 precursor (NID-2) (Osteonidogen). 787 o Nidogen-2 precursor (NID-2) (Osteonidogen) 787 787 o nidogen 2 (osteonidogen) 787	CAA57709.1 nidogen
788 787 785 785	AAA57261.1 nidogen
787	NP_031387.1 nidoger
785	4112
711	XP_051712.2 similar
	AAH35608.1 Similar

Ŋ

_

٦ Ω $\stackrel{\sim}{\sim}$

WO 2005/000335 PCT/US2004/017322

206 733 209 NP_001863.1 plasma carboxypeptidase B2 isoform a preproprotein; thrombin-activatable fibrinolysis inhibitor; thrombin-activable fibrinolysis inhibitor; carboxypeptidase U; carboxypeptidase B-like protein; procarboxypeptidase U; procarboxypeptidase R; plasma procarboxypeptidase B NP_078929.1 hypothetical protein FLJ13902 hypothetical protein FLJ13902 AAH14298.1 F:2.13 (5to11) F:2.13 (5to 19) NM_025774 Mm.31643 NM_019775 Mm.24242 NP 080050.1 NP_062749.1

9E-66 3E-60 4E-99 1E-164 7E-62 6E-99 4E-54 SE-53 1E-167 1E-167 578 226 587 237 585 360 359 245 733 NP_057497.2 plasma carboxypeptidase B2 isoform b; thrombin-activable fibrinolysis inhibitor; thrombinctivatable fibrinolysis inhibitor; carboxypeptidase U; carboxypeptidase B-like protein; procarboxypeptidase U; procarboxypeptidase R; plasma procarboxypeptidase B NP_113600.1 GABA(A) receptor-associated protein like 1; early estrogen-regulated protein Carboxypeptidase B precursor (Pancreas-specific protein) (PASP) GABA(A) receptors associated protein like Chain R, Human II-10 IL-10r1 Complex interleukin 10 receptor, alpha precursor BAA90475.1 carboxypeptidase B-like protein AAH07057.1 carboxypeptidase B2 (plasma) procarboxypeptidase B NP_115957.1 CAA12163.1 NP_001549 P15086 117VR F:2.12 (7to19) (F:2.12 (YtoO) NM_020590 Mm.14638 NM_008348 Mm.26658 NP 032374.1 065615. £

15

	pdbl1KOT	Chain A. Solution Structure Of Human Gaba Receptor Associated Protein Gabarap	216	6E-57
	NP 009209.1	GABA(A) receptor-associated protein; GABA(A)-receptor-associated protein	216	6E-57
NM_011375 Mm.38248 F:2.12 (5to19)		NP_003887.1 sialyltransferase 9 (CMP-NeuAc.lactosylceramide alpha-2,3-sialyltransferase; GM3 synthase);	628	1E-179
NP 035505.1		ganglioside G(M3) Synthase		
	AAD14634.1	CMP-NeuAc.lactosylceramide alpha-2,3-	979	1E-178
	NP_006270.1	sialyltransferase 6 isoform j; Gal beta-1,3(4)GlcNAc alpha-2,3 sialyltransferase; CMP-N-	213	1E-53
		acetylneuraminate-beta-1,4-galactoside alpha-2,3-sialyltransferase; alpha-2,3-sialyltransferase II;		
		alpha 2,3-sialyltransferase III		
	AAL14347.1	Gal beta 1,3(4) GlcNAc alpha 2,3-sialyltransferase	202	2E-50
NM_013563 Mm.2923 F:2.11 (YtoO)	O) NP_000197	interleukin 2 receptor, gamma chain, precursor; Interleukin-2 receptor, gamma; common cytokine	488	1E-138
NP 038591.1		receptor gamma chain; CD132		
NM_021291 Mm.45874 F:2.11 (YtoM)	M) NP_055085	solute carrier family 7 (cationic amino acid transporter, y+ system), member 9; solute carrier	754	0
NP_067266.1		family 7, member 9, solute carrier family 7 (cationic amino acid, transporter, y+ system), member		
		6		
	CAB54003	glycoprotein-associated amino acid transporter lib0,+AT1	751	0
	NP_003477	solute carrier family 7 (cationic amino acid transporter, y^+ system), member 5; Membrane protein	347	3E-95
		E16; Solute carrier family 7, member 5; 4F2 light chain		
	AAC61479.	amino acid transporter E16	347	3E-95
	BAB70708	sodium-independent neutral amino acid transporter L.AT1	346	7E-95
	AAH39692	Similar to solute carrier family 7 (cationic amino acid transporter, y+ system), member 5	346	7E-95
	BAA75746	4F2 light chain	346	7E-95
	CAD62619	unnamed protein product	345	2E-94
	Q9UM01	Y+L amino acid transporter 1 (y(+)L-type amino acid transporter 1) (y+LAT-1) (Y+LAT1)	345	2E-94
		(Monocyte amino acid permease 2) (MOP-2).		
	NP_055146	solute carrier family 7, (cationic amino acid transporter, y+ system) member 11; cystine/glutamate	344	2E-94
-		transporter		

WO 2005/000335 PCT/US2004/017322

-				215		
			NP 003973	solute carrier family 7 (cationic anno acid transporter, y+ system), member 7	343	5E-94
			NP 003974	solute carrier family 7 (cationic amino acid transporter, y+ system), member 6	337	SE-92
			Q9UHIS	Large neutral amino acids transporter small subunit 2 (L-type amino acid transporter 2) (hLAT2).	328	2E-89
			BAB40574	cystine/glutamate exchanger	328	2E-89
			NP 036376	solute carrier family 7 (cationic amino acid transporter, y+ system), member 8	326	4E-89
			NP 062823	solute carrier family 7, member 10; asc-type amino acid transporter 1	323	4E-88
			BAA95120	y+L amino acid transporter-1	322	1E-87
			AAF05695	L amino acid transporter-2; LAT-2	314	3E-85
			CAD62616	unnamed protein product	210	6E-54
			CAD10393	amino acid transporter	209	1E-53
			NP 620172	amino acid transporter XAT2	207	3E-53
NM_010016 Mm.20236	0236	F:2.11 (7to11)	NP_000565.1	000565.1 decay accelerating factor for complement (CD55, Cromer blood group system); Decay-	367	1E-101
NP_034146.1				accelerating factor of		
		4	P08174	Complement decay-accelerating factor precursor (CD55 antigen)	365	1E-101
			AAA52167.1	decay-accelerating factor precursor	364	1E-100
			AAB48622.1	decay-acceleration factor	363	1E-100
			A26359	decay-accelerating factor, splice form 1 precursor	355	1E-97
i 			23200413	Chain R, Structural Model Of Human Decay-Accelerating Factor Bound To Echovirus 7 From	308	8E-64
				Cryo-Electron Microscopy		
			AAL25833.1	decay-accelerating factor 1 ab	243	6E-64
			AAL25835.1	decay-accelerating factor 4ab	243	6E-64
			AAL25834.1	decay-accelerating factor 3	242	7E-64
L16846 Mm.16596 AAA37327.1	6596	F:2.11 (7to19)	NP_001722.1	001722.1 B-cell translocation protein 1	348	2E-94
			NP_006754.J	_006754.1 BTG family, member 2; B-cell translocation gene 2 (pheochromacytoma cell-3); B-cell	211	3E-53
		T		Transionation gene 2		

okD protein 1209 0	ed protein,	1202	1202 1196 893	1202 1196 893	1202 1196 893 770	1202 1196 893 770	1202 1196 893 770 759	1202 1196 893 770 759	1202 1196 893 770 759	1202 1196 893 770 759 759	1202 1196 893 770 759 759 758	1202 1196 893 770 759 758 758 756	1202 1196 893 770 759 758 758 754	1202 1196 893 770 759 759 754 754	1202 1196 893 770 759 758 754 754 753	1202 1196 893 770 759 758 754 754 750	1202 1196 893 770 759 754 754 750 750	1202 1196 893 759 759 754 754 750 750 750
005338.1 heat shock 70kDa protein 5 (glucose-regulated protein, 78kDa); BiP; heat shock 70kD protein for contents of colleges-regulated protein.) (Bideose-regulated protein	(Bittonsoringuitation protein	Tmmunoglobulin heavy chai	9 (glucose-regulated protein, 70kD), meat-shork 70kD protein-7 (glucose-regulated protein, 70kD) BiP protein dnaK-type molecular chaperone HSPA5 precursor similar to 78 kDa glucose-regulated protein precursor (GRP 78) (Immunoglobulin heavy chain binding protein) (BIP) (Endoplasmic reticulum lumenal Ca2+ binding protein grp78)	F13605.1 BiP protein 4821 dnaK-type molecular chaperone HSPA5 precursor 688941.1 similar to 78 kDa glucose-regulated protein precursor (GRP 78) (Immunoglobulin heavy chain binding protein) (BIP) (Endoplasmic reticulum lumenal Ca2+ binding protein grp78) 606588.1 heat shock 70kDa protein 8 isoform 1; heat shock cognate protein, 71-kDa; heat shock 70kd	Gillicose-regulated protein, 70kD). Inter-silven 70kD protein-3 (gillicose-regulated protein). BiP protein dnaK-type molecular chaperone HSPA5 precursor similar to 78 kDa glucose-regulated protein precursor (GRP 78) (Immunoglobulin heavy chain binding protein) (BIP) (Endoplasmic reticulum lumenal Ca2+ binding protein grp78) heat shock 70kDa protein 8 isoform 1; heat shock cognate protein, 71-kDa; heat shock 70kd protein 10; heat shock cognate protein 54; constitutive heat shock protein 70; lipopolysaccharide-	Immunoglobulin heavy chai	AAF13605.1 BiP protein A29821 dnaK-type molecular chaperone HSPA5 precursor XP_088941.1 similar to 78 kDa glucose-regulated protein precursor (GRP 78) (Immunoglobulin heavy chain binding protein) (BIP) (Endoplasmic reticulum lumenal Ca2+ binding protein grp78) NP_06588.1 heat shock 70kDa protein 8 isoform 1; heat shock cognate protein, 71-kDa; heat shock 70kd protein 10; heat shock cognate protein 1 BO8107 Heat shock 70kDa protein 1 (HSP70-1) (HSP70-1/HSP70-2) NP_065336.2 heat shock 70kDa protein 1A; heat shock 70kD protein 1A; heat shock 70kD protein 1A; heat shock 70kDa protein; dnaK-	Immunoglobulin heavy chai iding protein grp78) in, 71-kDa; heat shock 70k rotein 70; lipopolysaccharide	Immunoglobulin heavy chai ding protein gp78) sin, 71-kDa; heat shock 70k rotein 70; lipopolysaccharide	Immunoglobulin heavy chai in, 71-kDa; heat shock 70k rotein 70; lipopolysaccharide shock-induced protein; dnak	Immunoglobulin heavy chai ding protein grp78) sin, 71-kDa; heat shock 70k rotein 70; lipopolysaccharide shock-induced protein; dnak	Immunoglobulin heavy chai ding protein grp78) sin, 71-kDa; heat shock 70k rotein 70; lipopolysaccharide shock-induced protein; dnak	Immunoglobulin heavy chaiding protein grp78) in, 71-kDa; heat shock 70k rotein 70; lipopolysaccharide shock-induced protein; dnak shock-induced protein; dnak	Immunoglobulin heavy chai ding protein grp78) in, 71-kDa; heat shock 70k rotein 70; lipopolysaccharide shock-induced protein; dnak	Immunoglobulin heavy chaiding protein grp78) in, 71-kDa; heat shock 70k rotein 70; lipopolysaccharide shock-induced protein; dnak shock-induced protein; dnak	ding protein grp78) yin, 71-kDa; heat shock 70k rotein 70; lipopolysaccharide shock-induced protein; dnak ock 70kD protein-2	78kD) 78kD) 78kD) 78kD) 78kD) 78kD) 78kD) 78kD) 78kD) 78kD) 78kD) 78kD) 78kD) 78kDa glucose-regulated protein precursor (GRP 78) (Immunoglobulin heavy chain binding protein) (BIP) (Endoplasmic reticulum lumenal Ca2+ binding protein gpp78) 78kDa glucose-regulated protein precursor (GRP 78) (Immunoglobulin heavy chain binding protein) (BIP) (Endoplasmic reticulum lumenal Ca2+ binding protein gpp78) 78kD 70kDa protein (J. InS-associated protein 1; heat shock rognate protein, 71-kDa; heat shock 70kDa protein 14; heat shock 70kDa protein 14; heat shock 70kDa protein 14; heat shock 70kDa protein 18, heat shock 70kDa protein 18, heat shock 70kDa protein 18, heat shock 70kDa protein 19, heat shock 70kDa protein 1-like 73527.1 heat shock 70kDa protein 1-like 73527.1 heat shock 70kDa protein 1-like 73527.1 heat shock 70kDa protein 1-like 73537.1 heat shock 70kDa protein 1-like, Heat-shock 70kD protein-like-1; heat shock 70kDa protein 1-like, Heat-shock 70kDa protein 1-like, Heat-shock 70kDa protein 1-like, Heat-shock 70kDa protein 1-like, Heat-shock 70kDa protein 1-like
5 (glucose-regulated protein, 78kD); Heat-shock 70kD protein-5 (glucose-regulated protein, 78kD)			iRP 78) (Immunoglobulū	BiP protein dnaK-type molecular chaperone HSPA5 precursor similar to 78 kDa glucose-regulated protein precursor (GRP 78) (Immunoglobulin he binding protein) (BIP) (Endoplasmic reticulum lumenal Ca2+ binding protein grp78)	RP 78) (Immunoglobuli Ca2+ binding protein gr ate protein, 71-kDa; hea	IRP 78) (Immunoglobulii Ca2+ binding protein gr nate protein, 71-kDa; hea	iRP 78) (Immunoglobulii Ca2+ binding protein gri rate protein, 71-kDa; hea t shock protein 70; lipopo	RP 78) (Immunoglobulio Ca2+ binding protein gr. aate protein, 71-kDa; hea t shock protein 70; lipopo 170-2) IA; heat shock-induced p	RP 78) (Immunoglobulio Ca2+ binding protein grg rate protein, 71-kDa; heat shock protein 70; lipopo 70-2) IA; heat shock-induced p	RP 78) (Immunoglobulio Ca2+ binding protein gr. late protein, 71-kDa; hea t shock protein 70; lipopo 10-2) 1A; heat shock-induced p	dnaK-type molecular chaperone HSPA5 precursor similar to 78 kDa glucose-regulated protein precursor (GRP 78) (Immunoglobulin hubinding protein) (BIP) (Endoplasmic reticulum lumenal Ca2+ binding protein grp78 heat shock 70kDa protein 8 isoform 1; heat shock cognate protein, 71-kDa; heat spotein 10; heat shock cognate protein 54; constitutive heat shock protein 70; lipopolys associated protein 1; LPS-associated protein 1 Heat shock 70kDa protein 1 (HSP70-1) (HSP70-1/HSP70-2) heat shock 70kDa protein 1A; heat shock 70kD protein 1B; heat shock 70kDa protein 1B; heat shock 70kDa protein 1B heat shock 70kDa protein 2; Heat-shock 70kDa protein-2	IRP 78) (Immunoglobuliu Ca2+ binding protein grg tate protein, 71-kDa; hea t shock protein 70; lipopo IA; heat shock-induced p IB Heat-shock 70kD protein	IRP 78) (Immunoglobuliu Ca2+ binding protein grigate protein, 71-kDa; heat shock protein 70; lipopo 14; heat shock-induced protein 18 Heat-shock 70kD protein Heat-shock 70kD protein	RP 78) (Immunoglobuliu Ca2+ binding protein grg late protein, 71-kDa; hea t shock protein 70; lipopo 10-2) IA; heat shock-induced p Heat-shock 70kD proteiu Heat-shock 70kD proteiu	iRP 78) (Immunoglobulii Ca2+ binding protein griute protein, 71-kDa; heatshock protein 70; lipopo 14; heat shock-induced protein 1B Heat-shock 70kD protein 1B	RP 78) (Immunoglobuliu Ca2+ binding protein grg ate protein, 71-kDa; hea tshock protein 70; lipopo 1A; heat shock-induced p HB Heat-shock 70kD protein Heat-shock 70kD protein	IRP 78) (Immunoglobuliu Ca2+ binding protein griute protein, 71-kDa; heat shock protein 70; lipopo IA; heat shock-induced protein IB Heat-shock 70kD protein IB	IRP 78) (Immunoglobuliu Ca2+ binding protein grigate protein, 71-kDa; hea tshock protein 70; lipopo 1A; heat shock-induced p IB Heat-shock 70kD protein IOM) in-like-1; heat shock 70k
			precursor rin precursor (GRP 78	precursor in precursor (GRP 78 culum lumenal Ca2+1	precursor in precursor (GRP 78 culum lumenal Ca2+1 leat shock cognate pr	precursor in precursor (GRP 78 culum lumenal Ca2+1 leat shock cognate pr constitutive heat shock	in precursor (GRP 78 culum lumenal Ca2+1 leat shock cognate proconstitutive heat shock cin 1 HSP70-1/HSP70-2)	in precursor (GRP 78 culum lumenal Ca2+1 leat shock cognate proconstitutive heat shock cin 1 HSP70-1/HSP70-2) TOkD protein 1A; heat shock cognate proconstitution of the shock cognate proconstitution of the shock constitution of the shock cognition of th	in precursor culum lumenal Ca2+1 teat shock cognate pr constitutive heat shock ein 1 HSP70-1/HSP70-2)	rin precursor (GRP 78 culum lumenal Ca2+1 leat shock cognate proconstitutive heat shock cin 1 HSP70-1/HSP70-2) 70kD protein 1B; heat shock read the shock cognate proconstitutive heat shock constitutive heat shock constitutive heat shock read 1	in precursor culum lumenal Ca2+1 teat shock cognate pr constitutive heat shocl ein 1 HSP70-1/HSP70-2) 70kD protein 1B 0kD protein 1B	rin precursor (GRP 78 culum lumenal Ca2+1 leat shock cognate proconstitutive heat shock cin 1 HSP70-1/HSP70-2) 70kD protein 1B 70kD protein 1B	in precursor culum lumenal Ca2+1 teat shock cognate pr constitutive heat shocl ein 1 HSP70-1/HSP70-2) 70kD protein 1B 0kD protein 2; Heat-s	in precursor culum lumenal Ca2+1 teat shock cognate pr constitutive heat shocl cin 1 HSP70-1/HSP70-2) 70kD protein 1A; hear-10kD protein 1B OkD protein 2; Heat-1	in precursor culum lumenal Ca2+1 teat shock cognate pr constitutive heat shocl ein 1 HSP70-1/HSP70-2) 70kD protein 1B 0kD protein 2; Heat-	in precursor culum lumenal Ca2+1 teat shock cognate pr constitutive heat shocl cin 1 HSP70-1/HSP70-2) 70kD protein 1A; heat-shocl 70kD protein 13: Heat-shocl	in precursor culum lumenal Ca2+1 teat shock cognate pr constitutive heat shocl cin 1 HSP70-1/HSP70-2) 70kD protein 1A; heat-sock of the protein 1B OkD protein 1B OkD protein 2; Heat-sock of the protein 2; Heat-sock of the protein 2; Heat-sock of the protein 3; Heat-sock of the protein 3; Heat-sock of the protein 4; Heat-sock of the protein 5; Heat-sock of the protein 6; Heat-sock of the protein 6; Heat-sock of the protein 7; Heat-sock of the protein 1; Heat-sock of the protein 1; Heat-sock of the protein 1; Heat-sock of the protein 2; Heat-sock of the protein 2; Heat-sock of the protein 3; Heat-sock of the protein 3; Heat-sock of the protein 4; Heat-sock of the protein 5; Heat-sock of the protein 6; Heat-sock of the protein 7; Heat-sock of the protein 7; Heat-sock of the protein 6; Heat-sock of	ein precursor (GRP 78 reulum lumenal Ca2+1 leat shock cognate proconstitutive heat shock constitutive heat shock ein 1 TOkD protein 1B OkD protein 1B OkD protein 2; Heat-shock reat-shock roke protein 2; Heat-shock roke protein 2; Heat-shock roke protein 1 reat-shock roke protein 1 reat-shock roke protein 1 reat-shock roke protein-1 reat-shock roke roke roke roke roke roke roke rok
tein, 78kD); Heat-s'		0 7 66644	perone HSPA5 prec	perone HSPA5 prec s-regulated protein r indoplasmic reticulu	perone HSPA5 precedured protein protei	perone HSPA5 precargonated protein pracagonated protein pracagonated protein 1; heat n 8 isoform 1; heat protein 54; com	perone HSPA5 precedulated protein pracegulated protein pracedoplasmic reticulum 8 isoform 1; heat prate protein 54; compassociated protein in 1 (HSP70.1) (HS	perone HSPA5 precargulated protein pracagnated protein pracagnated protein 1; heat a sosociated protein 54; constant (HSP70.1) (HS in 1 (HSP70.1) (HS in 1 (HSP70.1))	perone HSPA5 precarange are gulated protein produced protein produced protein 54; compare protein 54; compare protein 54; compare protein prot	perone HSPA5 precargulated protein randoplasmic reticulum 8 isoform 1; heat mate protein 54; com 1.4 heat shock 701 (HSP70-1) (HSP70-1) (HSP70-1) (HSP70-1) (HSP70-1) (HSP70-1) (HSP70-1)	perone HSPA5 precedulated protein pracegulated protein pracedoplasmic reticulum 8 isoform 1; heat mate protein 54; compassociated protein in 1 (HSP70.1) (HS 11A; heat shock 70) e HSP70-1	perone HSPA5 precargulated protein paregulated protein paregulated protein paregulate protein 54; compares protein 54; compares protein 54; compares protein paregulates protein paregulat	perone HSPA5 precedulated protein pracegulated protein pracedoplasmic reticulum 8 isoform 1; heat mate protein 54; compassociated protein in 1 (HSP70.1) (HS not shock 70) e HSP70.1 1 1 B; heat shock 70 at 1 b; heat shock 70 ce HSP70.1 2 heat shock 70 ce HSP70.1 2 heat shock 70 ce HSP70.1	perone HSPA5 precargulated protein paregulated protein pandoplasmic reticulum 8 isoform 1; heat massociated protein 54; control (HSP70.1) (HS n. 1A; heat shock 70 n. 1B; heat sh	perone HSPA5 precestregulated protein paragraph of protein particulur in 8 isoform 1; heat mate protein 54; compare protein 54; compare protein 54; compare protein in 1 (HSP70.1) (HS nat shock 70) and 1, heat shock 70] a	perone HSPA5 precargulated protein paregulated protein paregulated protein I; heat n 8 isoform 1; heat presence of the protein 54; concarsociated protein in 1 (HSP70.1) (HS n 1A; heat shock 70l n 1B; heat shock 70l n 1B; heat shock 70l n 2; heat shock 70l n 1B;	perone HSPA5 precestregulated protein paragraph of the protein I; heat not in 1 (HSP70.1) (HS) associated protein in 1 (HSP70.1) (HS) associated protein in 1 (HSP70.1) (HS) heat shock 70ll e HSP70.1 1 1 B; heat shock 70ll of 1.2; heat shock 70ll of 2; heat shock 70ll of 1.2; heat shock 70ll o	perone HSPA5 prec -regulated protein I adoplasmic reticulu n 8 isoform 1; heat n 8 isoform 1; heat -associated protein in 1 (HSP70.1) (HS) 11A; heat shock 70l 11B; heat shock 70l 12; heat shock 70l 12; heat shock 70l 12; heat shock 70l 12; heat shock 70l 11k; GC:33922) GC:33922) kDa protein 1-HOW kDa protein 1-HOW to 11ike; Heat-shock in 1-like; in 1-like
hock 70kDa protein 1cose-regulated prote)	rotein	•	dnaK-type molecular chaperone HSPA5 precursor similar to 78 kDa glucose-regulated protein precu	-type molecular chap ir to 78 kDa glucose- ng protein) (BIP) (En	-type molecular chap rr to 78 kDa glucose- ng protein) (BIP) (En	-type molecular chap tr to 78 kDa glucose- ng protein) (BIP) (En shock 70kDa protein in 10; heat shock cogn	dnaK-type molecular chaperone HSPA5 precursor similar to 78 kDa glucose-regulated protein precursor (GRP 7 binding protein) (BIP) (Endoplasmic reticulum lumenal Ca2+heat shock 70kDa protein 8 isoform 1; heat shock cognate protein 10; heat shock cognate protein 54; constitutive heat shocassociated protein 1; LPS-associated protein 1 Heat shock 70 kDa protein 1 (HSP70.1) (HSP70-1/HSP70-2)	rype molecular chap in to 78 kDa glucose- ng protein) (BIP) (En shock 70kDa protein in 10; heat shock cogn iated protein 1; LPS- ishock 70 kDa protein ishock 70kDa protein	dnaK-type molecular chaperone HS similar to 78 kDa glucose-regulated binding protein) (BIP) (Endoplasmi heat shock 70kDa protein 8 isofort associated protein 1; LPS-associated Heat shock 70 kDa protein 1 (HSP7) heat shock 70kDa protein 1A; heat type molecular chaperone HSP70-1	dnaK-type molecular chaperone HSPA5 precursor similar to 78 kDa glucose-regulated protein precursor (GRI binding protein) (BIP) (Endoplasmic reticulum lumenal Caheat shock 70kDa protein 8 isoform 1; heat shock cognate protein 54; constitutive heat slassociated protein 1; LPS-associated protein 1 Heat shock 70 kDa protein 1 (HSP70.1) (HSP70-1/HSP70 heat shock 70kDa protein 1A; heat shock 70kD protein 1A type molecular chaperone HSP70-1	-type molecular chap rr to 78 kDa glucoseng protein) (BIP) (En shock 70kDa protein iated protein 1; LPS-shock 70kDa protein nolecular chaperone thock 70kDa protein indock 70kDa protein	A29821 dnaK-type molecular chapcrone HSI XP_088941.1 similar to 78 kDa glucose-regulated binding protein) (BIP) (Endoplasmi NP_006588.1 heat shock 70kDa protein 8 isoform protein 10; heat shock cognate protein associated protein 1; LPS-associated P08107 Heat shock 70kDa protein 1A; heat shock 70kDa protein 1B; heat shock 70kDa protein 1; heat sh	rt to 78 kDa glucoseng protein) (BIP) (En ni pock 70kDa protein) in 10; heat shock cogniated protein 1; LPS-shock 70kDa protein hock 70kDa protein nolecular chaperone shock 70kDa protein hock 70kDa protein own (protein for MG hock protein for MG hock protein	duak-type molecular chaperone HSPA5 psimilar to 78 kDa glucose-regulated protei binding protein) (BIP) (Endoplasmic retic heat shock 70kDa protein 8 isoform 1; hprotein 10; heat shock cognate protein 54; cassociated protein theat shock 70kDa protein 1 (HSP70.1) (heat shock 70kDa protein 1A; heat shock 70kDa protein 1B; heat shock protein	A29821 dnaK-type molecular chaperone XP_088941.1 similar to 78 kDa glucose-regul binding protein) (BIP) (Endople NP_006588.1 heat shock 70kDa protein 8 iso protein 10; heat shock cognate passociated protein 1; LPS-assoc P08107 Heat shock 70kDa protein 1 (H NP_005336.2 heat shock 70kDa protein 1A; h type molecular chaperone HSP NP_068814.2 heat shock 70kDa protein 1B; h NP_068814.2 heat shock 70kDa protein 2; heat shock 70kDa protein 1B; h AAD11466.1 heat shock protein A29160 dnaK-type molecular chaperone XP_175177.1 heat shock 70kD protein 1-like	dnaK-type molecular chaperone HS similar to 78 kDa glucose-regulated binding protein) (BIP) (Endoplasmi heat shock 70kDa protein 8 isoform protein 10; heat shock cognate protein associated Protein 1; LPS-associated Heat shock 70kDa protein 1A; heat shock 70kDa protein 1B; heat sheat shock 70kDa protein 1B; heat sheat shock 70kDa protein 1B; heat sheat shock protein for MGC:33922) theat shock protein HSP-heat shock protein HSP-heat shock protein HSP-heat shock 70kD protein 1-like Heat shock protein 70 testis variant	dnak-type molecular chaperone HSPA5 precursor binding protein) (BIP) (Endoplasmic reticulum lumenal Ca2+ binding protein) (BIP) (Endoplasmic reticulum lumenal Ca2+ binding protein) (BIP) (Endoplasmic reticulum lumenal Ca2+ binding protein 10; heat shock 70kDa protein 8 isoform 1; heat shock cognate protein 154; constitutive heat shock 70kDa protein 14; LPS-associated protein 1 brat shock 70kDa protein 14; HSP70-1/HSP70-1/HSP70-2) beat shock 70kDa protein 18; heat shock 70kD protein 18; heat shock 70kDa protein 2; heat shock 70kDa protein 2; heat shock 70kDa protein 2; heat shock 70kDa protein 19; heat shock 70kDa protein 19; heat shock 70kDa protein 10; heat shock 70kDa prot	duak-type molecular chaperone H similar to 78 kDa glucose-regulate binding protein) (BIP) (Endoplasn heat shock 70kDa protein 8 isofo protein 10; heat shock cognate prot associated protein 1; LPS-associat Heat shock 70kDa protein 1 (HSI heat shock 70kDa protein 18; hea heat shock 70kDa protein 18; hea heat shock 70kDa protein 18; hea heat shock 70kDa protein 11; heat heat shock 70kDa protein 11; heat shock protein Heat shock protein heat shock protein similar to Heat shock 70 kDa prot heat shock 70kDa protein 1-like heat shock 70kDa protein 1-like similar to Heat shock 70 kDa prot heat shock 70kDa protein 1-like; I i; heat shock 70kDa protein 1-like; I
5 (gluco 78kD)	5.1 BiP protein				dnaK-ty 41.1 similar t binding 88.1 heat sho	dnaK-ty 41.1 similar t binding 88.1 heat sho protein	dnaK-ty 41.1 similar t binding 88.1 heat sho protein associat Heat sho	dnaK-ty 41.1 similar t binding 88.1 heat sho protein associat Heat sho 36.2 heat sho	dnaK-ty 41.1 similar t binding 88.1 heat sho protein associat Heat sho 16.2 heat sho type mo	dnaK-ty 41.1 similar t binding 88.1 heat sho protein associat Heat sho 190.2 heat sho 190.2 heat sho 190.37.1 heat sho	dnaK-ty 41.1 similar t binding 88.1 heat sho protein associat Heat sho 14.2 heat sho 14.2 heat sho	dnaK-fy 41.1 similar t binding 88.1 heat sho protein associat Heat sho 14.2 heat sho 14.2 heat sho	A29821 dnaK-type molecul XP_088941.1 similar to 78 kDa g binding protein) (B NP_006588.1 heat shock 70kDa associated protein P08107 Heat shock 70kDa NP_005336.2 heat shock 70kDa type molecular cha type molecular cha NP_068814.2 heat shock 70kDa NP_068814.2 heat shock 70kDa AAH36107.1 Unknown (protein AAD11466.1 heat shock protein	dnaK-ty 41.1 similar t binding 88.1 heat sho protein 1 associatt Heat sho 14.2 heat sho 14.2 heat sho 17.1 Unkmow 56.1 heat sho dnaK-ty	dnaK-ty 41.1 similar t binding 88.1 heat sho protein 1 associat Heat sho 14.2 heat sho 07.1 Unknow dnaK-ty dnaK-ty T7.1 heat sho	dnaK-ty dnakk-ty binding 88.1 heat sho protein 1 associatt Heat sho 14.2 heat sho 17.1 Unkmow dnaK-ty dnak-ty T7.1 heat sho	dnaK-ty 41.1 similar t binding 88.1 heat sho protein 1 associat Heat sho 14.2 heat sho 07.1 Unknow dnaK-ty dnaK-ty 77.1 heat sho 48.1 similar t	dnaK-ty dnakk-ty 41.1 similar t binding 88.1 heat sho protein 1 associat Heat sho 14.2 heat sho 97.1 Unknow dnaK-ty 77.1 heat sho 21.1 Heat sh 48.1 similar t 18.1 heat sh 118.1 heat sh
·	AAF13605.1	10000	A29821 XP_088941.1	A29821 XP_088941	A29821 XP_088941 NP_006588	A29821 XP_088941 NP_006588	A29821 XP_088941 NP_006588	A29821 XP_088941 NP_006588 P08107 NP_005336	A29821 XP_088941 NP_006588 P08107 NP_005336	A29821 XP_088941.1 NP_006588.1 P08107 NP_005336.2 NP_005337.1	A29821 XP_088941.1 NP_006588.1 P08107 NP_005336.2 NP_005337.1 NP_068814.2	A29821 XP_088941 NP_006588 NP_005336 NP_005337 NP_068814 AAH36107	A29821 XP_088941 NP_006588 NP_005336 NP_005337 NP_068814 AAH36107 AAD11466	A29821 XP_088941 NP_006588 NP_005336 NP_068814 AAH36107 AAD11466 A29160	A29821 XP_088941 NP_006588 NP_005336 NP_005337 NP_068814 AAH36107 AAD11466 A29160 XP_175177	A29821 XP_088941.1 NP_006588.1 NP_005336.2 NP_068814.2 AAH36107.1 AAH36107.1 AAD11466.1 A29160 XP_175177.1 BAA32521.1	A29821 XP_088941 NP_006588 P08107 NP_005337 NP_068814 AAH36107 AAD11466 A29160 XP_175177 XP_166348	A29821 XP_088941 NP_006588 NP_0065336 NP_068814 AAH36107 AAD11466 A29160 XP_175177 XP_176348 NP_166348 NP_065518
F:2.11 (7to19) NP	7	†																
1 (AIRILL)		_																
NM_022310 Mm.918 NP_071705.1																		

		i	21/		
		AAH35665.1	AAI135665.1 heat shock 70kDa protein 6 (HSP70B')	736	0
		NP_002146.1	_002146.1 heat shock 70kDa protein 6 (HSP70B'); heat shock 70kD protein 6 (HSP70B'); Heat-shock 70kD protein-6 (HSP70B')	735	0
		AAH07276.1	Similar to heat shock cognate 71-kd protein	682	0
		BAB18615.1	BAB18615.1 heat shock cognate protein 54	009	1E-171
		NP_004125.2	NP_004125.2 heat shock 70kDa protein 9B (mortalin-2); heat shock 70kD protein 9 (mortalin); mot-2; mthsp75; heat shock 70kD protein 9B (mortalin-2); Heat-shock 70kD protein-9 (mortalin)	574	1E-163
A K 0 0 4 6 5 4 Mm.86439 1 BAB23445.1	F.2.11 (5to7)	AAH29926.1	Similar to hypothetical protein FLJ13511	927	0
		AAC18917.1 F02569_2	F02569_2	397	1E-109
		NP_149014.1	NP_149014.1 hypothetical protein FLJ13511	335	3E-90
		AAF22843.1 7h3 protein	7h3 protein	249	1E-64
A K 0 0 9 5 6 3 Mm.28697 BAB26361.1	F:2.1 (5to19)	XP_045585.1	045585.1 similar to Protein KLAA1434	929	0.
		BAA92672.1	KIAA1434 protein	929	0
		BAA91994.1	unnamed protein product	444	1E-124
NM_011579 Mm.15793 NP 035709.1	F:2.1 (5to19)	NP_062558.1	NP_062558.1 hypothetical protein R30953_1	233	4E-60
NM_021394 Mm.116687 F.2.1 (5to19) NP_067369.1	F:2.1 (5to19)	CAC17634.2	CAC17634.2 dJ718J7.3.1 (novel protein similar to mouse tumour stroma and activated macrophage protein DLM-1, isoform 1)	320	4E-87

			210		
		NP_110403.1	P_110403.1 tumor stroma and activated macrophage protein DLM-1; chromosome 20 open reading frame 183	308	1E-83
NM_016702 Mm.7457	F:2.1 (5to19)	NP_000021.1		919	1E-176
NP 057911.1			peroxisonal, serine-pynuvate aminotransferase		
		BAA02632.1	alanine:glyoxylate aminotransferase	614	1E-175
		1704252A	Ala/glyoxylate aminotransferase	609	1E-174
		AAK30157.1	hepatic peroxysomal alanine:glyoxylate aminotransferase	564	1E-160
NM_013550 Mm.193557 F:2.09 (YtoO)	F:2.09 (YtoO)	NP_003529	H4 histone family, member A	162	8E-41
NP 038578.1					
A K 0 0 3 9 3 8 Mm.6671	F:2.09 (YtoM)	BAB47495	KIAA1866 protein	1197	0
BAB23084.1					
		XP 027658	similar to KLAA1866 protein	1186	0
NM_019571 Mm.31927	F:2.09 (YtoM)	NP_005714	tetraspan 5; tetraspan TM4SF; tetraspan NET-4; transmembrane 4 superfamily member 9;	999	1E-161
NP 062517.1			transmenthrane 4 superfamily, member 8; tetraspanin 5		
		A59261	tetraspan TSPAN-5	552	1E-157
		XP 030295	similar to RIKEN cDNA 2210021G21 gene [Mus musculus]	447	1E-125
		AAM94899	DC-TM4F2 precursor	338	3E-92
		NP 112189	tetraspanin similar to TM4SF9	337	6E-92
		AAH02920	Similar to transmembrane 4 superfamily member 9	299	2E-80
		AAH44244	Similar to hypothetical protein MGC30714	525	2E-59
		BAB15717	FLJ00016 protein	212	3E-54
NM_007509 Mm.10727	F:2.09 (7to19)	NP_001684.2	NP_001684.2 ATPase, H+ transporting, lysosonal 56/58kD, VI subunit B, isoform 2; vacuolar proton pump	1008	0
NP_031535.2			B isoform 2; endomembrane proton pump 58 kDa subunit; vacuolar ATP synthase subunit B,		
			brain isoform; V-ATPase B2 subunit; H(+)-transporting two-sector ATPase. 56/58kD subunit.		•
			12		

2

7

딘

 $\tilde{\alpha}$

	B44138	H+-exporting ATPase (EC 3.6.3.6) 56K chain, vacuolar, brain isoform	1007	0
	AAA58661.1	vacuolar H+-ATPase 56,000 subunit	1005	0
	AAH30640.1	AAH30640.1 ATPase, H+ transporting, lysosomal 56/58kD, V1 subunit B, isoform 2	1002	0
	AAH07309.1	AAH07309.1 Unknown (protein for IMAGE:3352651)	066	0
	NP_001683.2	NP_001683.2 ATPase, H+ transporting, lysosomal 56/58kD, V1 subunit B, isoform 1; ATPase, H+ transporting, lysosomal, beta polypeptide, 58kD; vacuolar proton pump, subunit 3; vacuolar ATP	898	0
		synthase subunit B, kidney isoform; V-ATPase B1 subunit; endomembrane proton pump 58 kDa		
		subunit; H(+)-transporting two-sector ATPase, 58kD subunit; H+-ATPase beta 1 subunit; ATPase, H+ transporting, lysosomal 56/58kD, V1 subunit B, isoform 1 (Renal tubular acidosis		
		with deafness)		
	P15313	Vacuolar ATP synthase subunit B, kidney isoform (V-ATPase B1 subunit) (Vacuolar proton pump B isoform 1) (Endomembrane proton pump 58 kDa subunit)	863	0
	AAA36498.1	AAA36498.1 proton pump 58 kDa subunit	863	0
	A39212	H+-transporting ATPase (EC 3.6.1.35), vacuolar	962	0
	AAD11943.1	AAD11943.1 H+-ATPase beta 1 subunit	788	0
	AAH35978.1	Unknown (protein for MGC:32642)	417	1E-115
NM_008932 Mm.195966 F:2.09 (7to19) NP_032958.1	NP_000940.1	NP_000940.1 prolactin receptor	338	1E-124
	AAL23915.1	prolactin receptor short isoform Ia	338	1E-124
	AAD49855.1	AAD49855.1 intermediate prolactin receptor isoform	338	1E-124

	337 1E-123	337 1E-92	230 7E-92	242 4E-90	230 1E-86	271 2E-72	230 8E-61	234 1E-60	234 1E-60		protein 9 1176 0		470 1E-125	256 2E-68	gprotein) 256 2E-68	256 2E-68	256 3E-68	253 1E-67	
220	prolactin receptor short isoform 1b	AAM11661.1 delta 4-SF1b truncated prolactin receptor	AAM18048.1 prolactin receptor delta 7/11	Chain A, The Xray Structure Of A Growth Hormone-Prolactin Receptor Complex	prolactin receptor - human	prolactin receptor isoform delta S1 precursor	AAM11660.1 delta 4-delta 7/11 truncated prolactin receptor	Similar to RIKEN cDNA 1110029A09 gene	BAC04633.1 unnamed protein product		004791.1 transmembrane 9 superfamily member 2; 76 kDa membrane protein; transmembrane protein 9	superfamily member 2	NP_055557.1 KIAA0255 gene product	transmembrane protein TM9SF3	XP_050993.1 similar to Transmembrane 9 superfamily protein member 3 precursor (SM-11044 binding protein) (EP70-P-iso)	SM-11044 binding protein	unnamed protein product	unnamed protein product	1 - 1 - 1 - 1 - 1 - 1 - 1 - 1 - 1 - 1 -
	AAL23914.1	AAM11661.1	AAM18048.1	pdb 1BP3	A57018	AAK32703.1	AAM11660.1	AAH36923.1	BAC04633.1		NP_004791.1		NP_055557.1	AAF98159.1	XP_050993.1	AAF21983.1	BAB55369.1	BAA91362.1	DAC11929 1
								F:2.09 (5to19) -			F:2.09 (5to19)							•	
										,									
	. ,							A K 0 0 3 9 5 0 Mm.36072 BAB23088.1		.,	A K 0 1 0 3 2 5 Mm. 5885	NP 542123.1							

20 Ŋ

WO 2005/000335 PCT/US2004/017322

				221		
			NP_006396.2	NP_006396.2 transmembrane 9 superfamily member 1; multispanning membrane protein (70kD);	251	4E-65
				transmembrane protein 9 superfamily member 1		
			015321	Transmembrane 9 superfamily protein member 1 precursor (hMP70)	247	5E-64
NM_011521 Mm.3815 NP 035651.1	n.3815	F:2.09 (5to19)	P31431	Syndecan-4 precursor (Amphiglycan) (SYND4) (Ryndocan core protein)	238	2E-62
			NP_002990.1	NP_002990.1 syndecan 4 (amphiglycan, ryudocan)	238	2E-62
NM_019437 Mm.7013	n.7013	F:2.09 (5to19)	NP_060809.2	NP_060809.2 hypothetical protein FLJ11149	308	3E-83
NP 062310.1						
			BAA92033.1	unnamed protein product	306	1E-82
NM_007811 Mm.42230	n.42230	F:2.08 (5to11)	NP_000774.2	NP_000774.2 cytochrome P450, subfamily XXVIA, polypeptide 1, isoform 1; P450, retinoic acid-inactivating,	901	0
NP 031837.1				1; retinoic acid-metabolizing cytochrome; retinoic acid 4-hydroxylase		
			043174	Cytochrome P45026 (Retinoic acid-metabolizing cytochrome) (P450RAI) (hP450RAI) (Retinoic	968	0
				acid 4-hydroxylase)		
			NP_476498.1	NP_476498.1 cytochrome P450, subfamily XXVIA, polypeptide 1, isoform 2; P450, retinoic acid-inactivating,	813	0
				1; retinoic acid-metabolizing cytochrome; retinoic acid 4-hydroxylase		
•			NP_063938.1	AP_063938.1 cytochrome P450 retinoid metabolizing protein	391	1E-107
NM_010324 Mm.19039	n.19039	F:2.08 (5to11)	S29028	aspartate transaminase (EC 2.6.1.1) (clone 8C7)	810	0
NP 034454.1						
			S13035	aspartate transaminase (EC 2.6.1.1)	6//	0
			NP_002070.1	aspartate aminotransferase 1; glutamic-oxaloacetic transaminase 1, soluble	179	0
			AAH00525.1	glutamic-oxaloacetic transaminase 2, mitochondrial (aspartate aminotransferase 2)	395	1E-109
NM 008364 Mm. 24771	n.24771	F:2.08 (5to19)	NP 002173.1	NP 002173.1 interleukin 1 receptor accessory protein isoform 1	1028	0

Ŋ

NP 032390 1					
		NP 608273.1	608273.1 interleukin 1 receptor accessory protein isoform 2	645	0
-					
		NP_059112.1	059112.1 interleukin 1 receptor accessory protein-like 2; interleukin 1 receptor 9; IL-1 receptor; X-linked	320	1E-85
			interleukin-1 receptor accessory protein-like 2; IL-1 receptor accessory protein-like 2		
		AAF59412.1	X-linked interleukin-1 receptor accessory protein-like 2	319	2E-85
NM_023580 M	NM_023580 Mm.133330 F:2.08 (5to19)	P21709	Ephrin type-A receptor 1 precursor (Tyrosine-protein kinase receptor EPH)	1646	0
NP 076069.1					
		A34076	protein-tyrosine kinase (EC 2.7.1.112) receptor type eph 1 precursor	1573	0
		NP_005223.1	005223.1 EphA1; eph tyrosine kinase 1 (erythropoietin-producing hepatoma amplified sequence; oncogene	1569	0
			EPH; ephrin receptor EphA1); eph tyrosine kinase 1 (erythropoietin-producing hepatoma		
			amplified sequence); ephrin receptor EphA1		
		S44280,	receptor tyrosine kinase eph	1179	0
		AAH37166.1	EphA2	874	0
		NP_004422.1	EphA2; ephrin receptor EphA2; epithelial cell receptor protein tyrosine kinase	867	С
		NP_004431.1	004431.1 EphA7; Hek11; ephrin receptor EphA7	790	0
		NP_004429.1	EphA4; Hek8; TYRO1 protein tyrosine kinase; ephrin receptor EphA4	778	To
		178843	receptor protein-tyrosine kinase - human (fragment)	773	0
		XP_046083.2	similar to Ephrin type-A receptor 5 precursor (Tyrosine-protein kinase receptor EHK-1) (Eph	770	0
			homology kinase-1) (Receptor protein-tyrosine kinase HEK7)		
	· · · · · · · · · · · · · · · · · · ·	P54756	Ephrin type-A receptor 5 precursor (Tyrosine-protein kinase receptor EHK-1) (Eph homology kinase-1) (Receptor protein-tyrosine kinase HEK7)	768	0
		P29320	Ephrin type-A receptor 3 precursor (Tyrosine-protein kinase receptor ETK1) (HEK4)	762	0
		NP_005224.2	005224.2 EphA3; Ephrin receptor EphA3 (human embryo kinase 1); eph-like tyrosine kinase 1 (human	759	0
		·	embryo kinase 1); ephrin receptor EphA3		
	-	NP_004432.1	004432.1 ephrin receptor EphB1 precursor; eph tyrosine kinase 2; ephrin receptor EphB1	708	0

کا

 Ξ

'

WO 2005/000335 PCT/US2004/017322

		223		
	NP_004434.2	004434.2 ephrin receptor EphB3 precursor; human embryo kinase 2; EPH-like tyrosine kinase 2; tyrosine-	702	0
	AAD02030.1	Eph-like receptor tyrosine kinasc hEphB1	701	0
	NP_065387.1	065387.1 ephrin receptor EphA8 precursor; ephrin type-A receptor 8 precursor; eph- and elk-related	669	0
		tyrosine kinase; tyrosylprotein kinase; tyrosine-protein kinase receptor eel; protein-tyrosine		
·	 	kinase; hydroxyaryl-protein kinase		
	P54753	Ephrin type-B receptor 3 precursor (Tyrosine-protein kinase receptor HEK-2)	969	0
	AAD02031.1	Eph-like receptor tyrosine kinase hEphB1b	691	0
	178842	receptor protein-tyrosine kinase - human (fragment)	687	0
	P29323	Ephrin type-B receptor 2 precursor (Tyrosine-protein kinase receptor EPH-3) (DRT) (Receptor	989	0
		protein-tyrosine kinase HEK5) (ERK)		
-	AAA99310.1	protein-tyrosine kinase	989	0
	NP_059145.1	ephrin receptor EphB2 isoform 1 precursor; developmentally-regulated eph-related tyrosine	989	0
		kinase; elk-related tyrosine kinase; eph tyrosine kinase 3		,
	BAA06506.1	tyrosine kinase precursor	685	0
	NP_004435.2	004435.2 ephrin receptor EphB4 precursor, Ephrin receptor EphB4 (hepatoma transmembrane kinase);	959	0
		Tyro11; ephrin receptor EphB4; hepatoma transmembrane kinase		
	 AAA20598.1	tyrosine kinase	959	0
	P54760	Ephrin type-B receptor 4 precursor (Tyrosine-protein kinase receptor HTK)	959	0
	AAB94627.1	Eph-like receptor tyrosine kinase hEphB1c	650	0
•	AAL14195.1	receptor protein tyrosine kinase variant EphB4v1	909	1E-173
	NP_004436.1	004436.1 ephrin receptor EphB6 precursor; tyrosine-protein kinase-defective receptor; ephrin type-B	598	1E-170
		i eceptor o		

2	
$^{\prime}$	

BAA03583.1 KIAA1459 protein 447 1E-125 AAD03088.1 Eph-family protein 447 1E-125 CAC10350.1 d174MI.1.1 (tyrosine kinase isosform 1) 389 1E-110 CAC10351.1 d174MI.1.2 (tyrosine kinase isosform 2) 389 1E-100 CAC10351.1 large erk kinase 381 1E-109 CAA65265 plateophilin 2a 614 1E-174 NP 004563. platophilin 2a 614 1E-174 NP 109589.2 phosphotriesterase-related 627 1E-179 AAK14923.1 HPHRP 627 1E-179 NP 003136.1 signal sequence receptor, beta (translocon-associaled protein beta) 339 5E-39 NP 000304.1 protein beta 1092 100	AAH04264.1	4AH04264.1		AAH04264.1 Similar to EphB4	594	1E-169
AAD03058.1 Eph-family protein 447 CAC10350.1 d174M1.1.1 (tyrosine kinase isosform 1) 399 CAC10351.1 d174M1.1.2 (tyrosine kinase isosform 2) 399 AS7174 protein-tyrosine kinase (BC 2.7.1.112) erk - human (fragment) 394 AAG43577.1 large erk kinase 351 CAA66265 plakophilin 2a 614 NP 004563. plakophilin 2 614 NP_109580.2 plosphotriesterase-related 614 AAK14923.1 HPHRP 627 AAK14923.1 Fignal sequence receptor, beta (translocon-associated protein beta) 339 NP_003304.1 signal sequence receptor, beta precursor; Signal sequence receptor, beta; translocon-associated protein beta 337 NP_000304.1 protein S (alpha); Protein S, alpha 1092			3AA95983.1	KIAA1459 protein	592	1E-168
CAC10350.1 dJ74M1.1.1 (tyrosine kinase isoform 1) 399 CAC10351.1 dJ74M1.1.2 (tyrosine kinase isosform 2) 399 A57174 protein-tyrosine kinase (EC 2.7.1.112) erk - human (fragment) 394 AAG43577.1 arge erk kinase 351 BAA03537.1 large erk kinase 351 CAA66265 plakophilin 2 635 NP 004563. plakophilin 2 635 NP 109589.2 phosphotriesterase-related 627 AAK14923.1 HPHRP 627 AAK14923.1 Fignal sequence receptor, beta (translocon-associated protein beta) 339 NP_00336.1 signal sequence receptor, beta (translocon-associated protein beta) 339 NP_000304.1 protein beta 1092 NP_000304.1 protein S (alpha); Protein S, alpha 1092		-	4AD03058.1	Eph-family protein	447	1E-125
CAC10351.1 d/74M1.1.2 (tyrosine kinase isosform 2) 399 A57174 protein-tyrosine kinase (EC 2.7.1.112) erk - human (fragment) 394 AAG43577.1 ephtrin receptor EPHA3 secreted form 385 BAA03537.1 large erk kinase 351 CAA66265 plakophilin 2 635 NP 004563. plakophilin 2 614 NP 109589.2 phosphotriesterase related 637 AAK14923.1 HPHRP 627 AAK14923.1 FPHRP 627 AAH00341.1 signal sequence receptor, beta (translooon-associated protein beta) 339 NP_000304.1 protein S (alpha); Protein S, alpha 1092			CAC10350.1	d174M1.1.1 (tyrosine kinase isoform 1)	399	1E-110
AAG43577.1 ephrin receptor EPHA3 secreted form BAA03537.1 large crk kinase CAA66265 plakophilin 2a NP_109589.2 plosphotriesterase related; resiniferatoxin-binding, phosphotriesterase-related AAK14923.1 HPHRP AAH00341.1 signal sequence receptor, beta (translocon-associated protein beta) NP_003136.1 signal sequence receptor, beta precursor; Signal sequence receptor, beta protein beta NP_000304.1 protein S (alpha); Protein S, alpha			ÇAC10351.1.	d174M1.1.2 (tyrosine kinase isosform 2)	399	1E-110
AAG43577.1 ephrin receptor EPHA3 secreted form 385 BAA03537.1 large erk kinase 351 CAA66265 plakophilin 2a 614 NP 109589.2 plakophilin 2 614 NP 109589.2 phosphotriesterase related 620 AAK14923.1 FFPHRP 627 AAK14923.1 FFPHRP 627 AAH00341.1 signal sequence receptor, beta (translocon-associated protein beta) 339 NP 000304.1 protein beta 1092 NP 000304.1 protein S (alpha); Protein S, alpha 1092		7	A57174	protein-tyrosine kinase (EC 2.7.1.112) erk - human (fragment)	394	1E-109
BAA03537.1 large crk kinase CAA66265 plakophilin 2a CAA66265 plakophilin 2a NP 004563. plakophilin 2 NP 109589.2 phosphotriesterase related; resiniferatoxin-binding, phosphotriesterase-related and the phosphotriesterase-related AAK14923.1 HPHRP AAK14923.1 HPHRP AAK14923.1 RPHRP AAK10341.1 signal sequence receptor, beta (translocon-associated protein beta) NP 003136.1 signal sequence receptor, beta precursor; Signal sequence receptor, beta; translocon-associated protein beta NP 000304.1 protein S (alpha); Protein S, alpha NP 000304.1 protein S (alpha); Protein S, alpha		7	AAG43577.1	ephrin receptor EPHA3 secreted form	385	1E-106
CAA66265 plakophilin 2a NP 004563. plakophilin 2 NP 019589.2 plosphotriesterase related; resiniferatoxin-binding, phosphotriesterase-related gene; 630 1 phosphotriesterase-related AAK14923.1 HPHRP AAH00341.1 signal sequence receptor, beta (translocon-associated protein beta) NP_063136.1 signal sequence receptor, beta precursor; Signal sequence receptor, beta; translocon-associated 337 protein beta NP_060304.1 protein S (alpha); Protein S, alpha			BAA03537.1	large erk kinase	351	4E-96
F:2.07 (YtoM) CAA66265 plakophilin 2a NP 004563. plakophilin 2 F:2.07 (7to19) NP_109589.2 phosphotriesterase related; resiniferatoxin-binding, phosphotriesterase-related phosphotriesterase-related AAK14923.1 HPHRP F:2.07 (7to19) AAK14923.1 HPHRP F:2.07 (7to19) AAH00341.1 signal sequence receptor, beta (translocon-associated protein beta protein beta NP_003136.1 signal sequence receptor, beta precursor; Signal sequence receptor, beta; translocon-associated 337 protein beta F:2.07 (5to19) NP_000304.1 protein S (alpha); Protein S, alpha						
F:2.07 (7to19) NP_109589.2 phosphotriesterase related; resiniferatoxin-binding, phosphotriesterase-related gene; 630 1 phosphotriesterase-related AAK14923.1 HPHRP F:2.07 (7to19) AAH00341.1 signal sequence receptor, beta (translocon-associated protein beta) NP_003136.1 signal sequence receptor, beta precursor; Signal sequence receptor, beta; translocon-associated 337 protein beta 1 F:2.07 (5to19) NP_000304.1 protein S (alpha); Protein S, alpha	Mm.196058 F:2.07		CAA66265	plakophilin 2a	635	
F:2.07 (7to19) NP_109589.2 phosphotriesterase related; resiniferatoxin-binding, phosphotriesterase-related gene; 630 1 phosphotriesterase-related AAK14923.1 FPHRP AAK14923.1 FPHRP F:2.07 (7to19) AAH00341.1 signal sequence receptor, beta (translocon-associated protein beta) P:2.07 (7to19) AAH00341.1 signal sequence receptor, beta precursor; Signal sequence receptor, beta; translocon-associated 337 protein beta P:2.07 (5to19) NP_000304.1 protein S (alpha); Protein S, alpha			NP 004563.	plakophilin 2	614	1E-174
F:2.07 (7to19) NP_109589.2 phosphotriesterase related; resiniferatoxin-binding, phosphotriesterase-related gene; 630 1 phosphotriesterase-related AAK14923.1 FPHRP AAK14923.1 FPHRP F:2.07 (7to19) AAH00341.1 signal sequence receptor, beta (translocon-associated protein beta) P:2.07 (7to19) NP_003136.1 signal sequence receptor, beta precursor; Signal sequence receptor, beta; translocon-associated 337 protein beta 1 F:2.07 (5to19) NP_000304.1 protein S (alpha); Protein S, alpha						
F.2.07 (7to19) AAH00341.1 signal sequence receptor, beta (translocon-associated protein beta) NP_003136.1 signal sequence receptor, beta precursor; Signal sequence receptor, beta; translocon-associated 337 protein beta 1 F.2.07 (5to19) NP_000304.1 protein S (alpha); Protein S, alpha	N.M_008961 Mm.2008 F:2.07 NP 032987.1		NP_109589.2	related; resiniferatoxin-binding,	l	•
F:2.07 (7to19) AAH00341.1 signal sequence receptor, beta (translocon-associated protein beta) 339 NP_003136.1 signal sequence receptor, beta precursor; Signal sequence receptor, beta; translocon-associated 337 protein beta 1 F:2.07 (5to19) NP_000304.1 protein S (alpha); Protein S, alpha 1092		,		HPHRP	627	•
F:2.07 (7to 19) AAH00341.1 signal sequence receptor, beta (translocon-associated protein beta) 339 NP_003136.1 signal sequence receptor, beta precursor; Signal sequence receptor, beta; translocon-associated 337 protein beta 1 F:2.07 (5to 19) NP_000304.1 protein S (alpha); Protein S, alpha 1092						
NP_003136.1 signal sequence receptor, beta precursor; Signal sequence receptor, beta; translocon-associated 337 2E-9 protein beta NP_000304.1 protein S (alpha); Protein S, alpha	NM_025448 Mm.7091 F:2.07 NP 079724.1		AAH00341.1	signal sequence receptor, beta (translocon-associated protein beta)	339	5E-93
F.2.07 (5to19) NP_000304.1 protein S (alpha); Protein S, alpha 1092			NP_003136.1	signal sequence receptor, beta precursor; Signal sequence receptor, beta; translocon-associated		2E-92
F:2.07 (Sto19) NP_000304.1 protein S (alpha); Protein S, alpha 1092				protein beta		
F:2.07 (5to19) NP_000304.1 protein S (alpha); Protein S, alpha 1092						
			NP_000304.1		1092	

10 12

Ŋ

			P07225	Vitamin K-dependent protein S precursor	1092	0
			AAA60180.1	protein S alpha	1092	0
			CAA31383.1	pre-protein S (AA -15 to 635)~ttg start	1082	0
			AAA60181.1	AAA60181.1 protein S precursor	1082	0
			NP_000811.1	growth arrest-specific 6; AXL stimulatory factor	548	1E-154
M12573	Mm.6388	F:2.07 (Sto 19)	NP_005336.2	NP_005336.2 heat shock 70kDa protein 1A; heat shock 70kD protein 1A; heat shock-induced protein; dnaK-	347	2E-94
AAA37863.1				type molecular chaperone HSP70-1		
			P08107	Heat shock 70 kDa protein 1 (HSP70.1) (HSP70-1/HSP70-2)	347	2E-94
	-		NP_005337.1	NP_005337.1 heat shock 70kDa protein 1B; heat shock 70kD protein 1B	345	6E-94
			A29160	A29160 dnaK-type molecular chaperone HSPA1L	341	2E-92
	•		XP_175177.1	heat shock 70kD protein 1-like	312	6E-84
			NP_005518.1	NP_005518.1 heat shock 70kDa protein 1-like; Heat-shock 70kD protein-like-1; heat shock 70kD protein-like	311	1E-83
				1; heat shock 70kD protein 1-like		
			BAA32521.1	Heat shock protein 70 testis variant	310	3E-83
			XP_166348.1	XP_166348.1 similar to Heat shock 70 kDa protein 1-HOM (HSP70-HOM)	310	3E-83
			AAH34483.1	heat shock 70kD protein 1-like	308	1E-82
			AAH07276.1	Similar to heat shock cognate 71-kd protein	301	1E-80
:	-		AAH15699.1	Unknown (protein for IMAGE:3906958)	301	1E-80
			NP_006588.1	NP_006588.1 heat shock 70kDa protein 8 isoform 1; heat shock cognate protein, 71-kDa; heat shock 70kd	301	1E-80
		-		protein 10; heat shock cognate protein 54; constitutive heat shock protein 70; lipopolysaccharide-		
	_			associated protein 1; LPS-associated protein 1		

10 10 15

	ı
O	1
a	1
S	ı
	1

			226	-	
	-	AAH08907.1	Similar to heat shock 70kD protein 8	301	1E-80
		NP 068814.2	068814.2 heat shock 70kDa protein 2; heat shock 70kD protein 2; Heat-shock 70kD protein-2	300	3E-80
		, , , , , , , , , , , , , , , , , , ,	Unknown (protein for MGC:33922)	300	3E-80
		AAD11466.1	heat shock protein	300	3E-80
		CAA36062.1	heat shock protein 70B' (AA 355-643)	285	1E-75
		XP .084070.5	similar to HEAT SHOCK 70 KD PROTEIN 6 (HEAT SHOCK 70 KD PROTEIN B)	285	1E-75
		H35665.1	AAH35665.1 heat shock 70kDa protein 6 (HSP70B')	285	1E-75
		NP_002146.1	002146.1 heat shock 70kDa protein 6 (HSP70B'); heat shock 70kD protein 6 (HSP70B'); Heat-shock 70kD	285	1E-75
			protein-6 (HSP70B')		
8 0 Mm.16106	F:2.07 (5to19)	P01028	Complement C4 precursor [Contains: C4A anaphylatoxin]	2587	0
NP 033910.1		241111	manufacture or Ivalitated	2586	0
		C4f1U			ľ
		NP_009224.1	009224.1 complement component 4A preproprotein; acidic C4; Rodgers form of C4; complement component 4S	2583	0
		CAB89302.1	dJ34F7.4 (complement component 4A)	2582	0
			This form of M. complement companient	2581	
		NP_000583.1	_000583.1 complement component 4.b preproprotem; Chido torm of C4, basic C4, comprement component.		
		AAB59537.1		2563	0
		AAA99717.1	complement C4B precursor	2465	0
		NP_000055.1	000055.1 complement component 3 precursor	624	1E-178
		AAA59651.1	complement component C4B	573	1E-163

	479 1E-134	1E-121	1E-119	1E-117	1E-100	1E-157	1E.153		1E-149		1E-148		1E-148	1E-148	1E-148		1E-130	1E-126	1E-116	6E-92		3 1E-89	- 1	5 5F.84
	479	436	430	421	366	553	5/10	5	525		523		523	522	521		462	448	415	.335		328	- 1	200
227	NP_001726.1 complement component 5	complement C4d	complement C4d variant	_	complement component C5	XP_003631.3 similar to ADP, ATP carrier protein, heart/skeletal muscle isoform T1 (ADP/ATP translocase 1)	(Adenine nucleotide translocator 1) (ANT1)		NP_001143.1 solute carrier family 25 (mitochondrial carrier; adenine nucleotide translocator), member 5; 2F1;	adenine nucleotide translocator 2 (fibroblast)	XP_114724.1 similar to ADP, ATP carrier protein, liver isoform T2 (ADP/ATP translocase 3) (Adenine	nucleotide translocator 3) (ANT 3)	ADP, ATP carrier protein T2	ADP/ATP carrier protein (adenine nucleotide translocator 2)	Similar to solute carrier family 25 (mitochondrial carrier; adenine nucleotide translocator),	member 5	ADP, ATP carrier protein (clone pHAT8) - human	ADP.ATP translocase	NP_112581.1 hypothetical protein DKFZp434N1235	1 similar to ADP,ATP carrier protein, fibroblast isoform (ADP/ATP translocase 2) (Adenine	nucleotide translocator 2) (ANT 2)	XP_070893.1 similar to ADP, ATP carrier protein, liver isoform T2 (ADP/ATP translocase 3) (Adenine	nucleotide translocator 3) (ANT 3)	17/0 1 (11/1/1/1/1/1/1/1/1/1/1/1/1/1/1/1/1/1
	NF_001726.1	1006226A	1006226B	AAC98380.1	AAA51856.1	XP_003631.3		NP_001142.1	NP_001143.		XP_114724.		A29132	AAB96347.1	AAH14775.1		B28116	AAA36749.1	NP_112581.	XP_065814.1		XP_070893.		
						228 F:2.07 (5to19)				 					5									
				-		 Mm.16228					_													
						U 2 7 3 1 5	AAC52837.1					-												- Contract of the Contract of

			1 (ANT 9) (ANT 9)		
			nucleotide translucator 2)	290	2E-78
		XP_095833.1 s	similar to ADF/ATF carrier protein		5
		XP_167013.1	similar to ADP,ATP carrier protein, fibroblast isoform (ADP/ATP translocase 2) (Adenine	255	1E-67
		-7.	nucleotide translocator 2) (ANT 2)	010	73 46
		XP_167333.1	similar to ADP, ATP carrier protein, fibroblast isoform (ADP/ATP translocase 2) (Ademine	718	QC-37
			nucleotide translocator 2) (ANT 2)	,	
		XP_063000.2	063000.2 similar to ADP/ATP carrier protein	196	6E-50
					- 1
NM_007860Mm.2774 F	F:2.06 (7to19)	P49895	Type I iodothyronine deiodinase (Type-I 5'deiodinase) (DIOI) (Type 1 DI) (5DI)	417	
		NP_000783.2	000783.2 thyroxine deiodinase type 1; 5DI; thyroxine deiodinase type I (selenoprotein)	409	1E-113
		AAH17955.1	Similar to deiodinase, iodothyronine, type I	207	4E-52
					ı
NM_0200001 Mm.117121 F:2.06 (5to19)	:2.06 (5to19)	NP_569708.1	569708.1 dendritic cell lectin b; blood dendritic cell antigen 2 protein	224	2E-58
NP 064385.1	`				
3 Mm.14798	F:2.06 (5to19)	NP_001008.1	ribosomal protein S13; 40S ribosomal protein S13	300	6E-82
NP 080809.1		A A C15854 1	ribosoma Inratein S13	225	3E-59
		AAC13034:1	HOUSONIAN PROCESS OF S		
7.3 Mm.20127	F:2.05 (YtoO)	BAA92054	unnamed protein product	598	1E-171
NP 203537.1			13. 1 1. 1 1. 1 1. 1 1. 1 1. 1 1. 1 1.	502	15 170
		NP_056330	keratin 23 isoform a; hyperacetylation-inducible type 1 keratin; keratin, type 1 cytoskejetal 23;		
		,	cytokeratin 23; type I intermediate filament cytokeratin; histone deacetylase inducible keratin 23		
		AAH28356	type I intermediate filament cytokeratin	593	L
		09C075	Keratin, type I cytoskeletal 23 (Cytokeratin 23) (K23) (CK 23).	591	11E-169

WO 2005/000335 PCT/US2004/017322

-			229	27.7	70 00
		T17294	hypothetical protein DKFZp434G032.1	4	/E-88
		NP 775320.	keratin 23 isoform b; hyperacetylation-inducible type I keratin; keratin, type I cytoskeletal 23;	321	2E-87
		l	cytokeratin 23: type I intermediate filament cytokeratin; histone deacetylase inducible keratin 23	3	
		237780	keratin 20. tvne I-like, cvtoskeletal	299	6E-81
		XP 049979	similar to Keratin, type I cytoskeletal 20 (Cytokeratin 20) (K20) (CK 20)	299	8E-81
		P08727	Keratin, type I cytoskeletal 19 (Cytokeratin 19) (K19) (CK 19).	287	3E-77
		NP_002267	keratin 19; keratin, type I cytoskeletal 19; keratin, type I, 40-kd; cytokeratin 19; 40-kDa keratin	n 287	3E-77
	-		intermediate filament precursor gene		
		BAC04534	unnamed protein product	287	3E-77
		NP 000413	keratin 17	287	3E-77
		KRHU9.	keratin 19, type I, cytoskeletal	286	7E-77
		NP 000214	keratin 12 (Meesmann corneal dystrophy); Keratin-12; keratin 12	283	4E-76
		NP 002266	keratin 15; keratin-15, basic; keratin-15, beta; type I cytoskeletal 15; cytokeratin 15	283	4E-76
		P19012	Keratin, type I cytoskeletal 15 (Cytokeratin 15) (K15) (CK 15).	283	4E-76
		NP 002265	keratin 13 isoform b; keratin, type I cytoskeletal 13; cytokeratin 13	281	2E-7
		NP 705694	keratin 13 isoform a; keratin, type I cytoskeletal 13; cytokeratin 13	281	2E-75
		KRHU3	keratin 13, type I, cytoskeletal, long splice form	281	2E-75
		AAA59460	keratin type 16	278	1B-74
		NP 005548	keratin 16; keratin, type I cytoskeletal 16; cytokeratin 16	278	2E-74
		JC4313	keratin 16, type I, cytoskeletal	278	2E-74
		KRHUE	keratin 14, type I, cytoskeletal	277	
		AAH02690	keratin 14 (epidermolysis bullosa simplex, Dowling-Meara, Koebner)	277	3E-74
		NP 000517	keratin 14; cytokeratin 14	277	_
		NP 003762	type I hair keratin 6; keratin, hair, acidic, 6	264	\Box
		AAH43581	Similar to keratin, hair, acidic, 6	264	
		CAA51914	cytokeratin 20	263	4E-70
		NP 002271	type I hair keratin 5; Ha-5; hard keratin, type I, 5	257	3E-68
		NP 061889	hypothetical protein FLJ20261	256	
		CAA76387		256	1
		Q92764		256	5 8E-68
		CAA62286	г	256	8E-68
		XP 039921	similar to keratin 17	253	3 SE-67
-		AAH34697	keratin 10 (epidermolytic hyperkeratosis; keratosis palmaris et plantaris)	252	9E-67
	The state of the s				

			230	0,0	7 17
		P13645	Keratin, type I cytoskeletal 10 (Cytokeratin 10) (K10) (CK 10).	707	70-97
		170564	similar to keratin 17	252	9E-67
		ì	tyme I hair keratin 3A: Ha-3I; hard keratin, type I,3I; keratin, hair, acidic,3A	251	2E-66
		000	Keratin tyne I cuticular HA3-I (Hair keratin, type I IHA3-I).	251	3E-66
		Т	keratin 10. type I. cytoskeletal	250	3E-66
		89	type I hair keratin 1; hard keratin, type I, 1; Ha-1; keratin, hair, acidic, 1	249	6E-66
		1223	Keratin. type I cuticular HA1 (Hair keratin, type I HA1).	249	7E-66
		7	Reratin, type I cuticular HA4 (Hair keratin, type I HA4).	248	1E-65
		NP 002270	tyne I hair keratin 3B; keratin, hair, acidic, 3B; Ha-3II; hard keratin, type I, 3II	248	2E-65
		7	keratin Ha1, type I, bair	247	4E-65
		CAA57956	hair keratin acidic 3-II	246	5E-65
		AAH41070	similar to keratin, hair, acidic, 4	246	6E-65
		ī	type I hair keratin 4; hard keratin, type I, 4	245	1E-64
		002269	type I hair keratin 2; Ha-2; hard keratin, type I, 2; keratin, hair, acidic,2	245	1E-64
			similar to RIKEN cDNA 4733401L19 [Mus musculus]	244	2E-64
		014532	Keratin, type I cuticular HA2 (Hair keratin, type I HA2).	- 244	2E-64
٠		CAA57179	hair type I acidic keratin	244	2E-64
		NP 000215	keratin 18	243	4E-64
		CAA82315	cytokeratin 9	243	7E-64
		CAA31377	cytokeratin 18 (424 AA)	243	7E-64
		NP 000217	keratin 9; Keratin-9	243	7E-64
			keratin Ha3-II, type I, hair - human	242	9E-64
		AAH00698	keratin 18	242	1E-63
		AAA59468	keratin-10	239	6E-63
		CAA76389	type I hair keratin 7	236	5E-62
		NP 000412	keratin 10; Keratin-10	236	5E-62
		076015	Keratin, type I cuticular HA8 (Hair keratin, type I HA8).	236	6E-62
		NP 006762	type I hair keratin 8	236	6E-62
		AAH09754	Similar to keratin 18	233	4E-61
		NP 003761	type I hair keratin 7	232	9E-61
		BAC03847	unnamed protein product	216	9E-56
A K 0 0 9 0 2 0 Mm.44194	4 F:2.05 (YtoM)	NP_004660	chloride intracellular channel 3	380	1E-105
BAB26030.2					

_

5

		+112	1000	27 25
	015247	Chloride intracellular channel protein 2 (XAP121).	677	35-00
	AAH22305	chloride intracellular channel 2	229	3E-60
	CAA73228	p64 bovine chloride channel-like protein	228	9E-60
	NP 001280	chloride intracellular channel 2	228	9E-60
	CAA03948	CLJC2	227	1E-59
	NP 001279	chloride intracellular channel 1; p64CLCP	226	2E-59
	AAC25675	nuclear chloride ion channel protein	225	5E-59
	Q9NZA1	Chloride intracellular channel protein 5	224	8E-59
	BAC11444	unnamed protein product	224	8E-59
	NP 058625	chloride intracellular channel 5	224	8E-59
	NP 039234	chloride intracellular channel 4; chloride intracellular channel 4 like	224	1E-58
	NP 444507	chloride intracellular channel 6; chloride channel form A	223	2E-58
	1K00A	Chain A, Crystal Structure Of A Soluble Form Of Clic1. An Intracellular Chloride Ion Channel	223	3E-58
	AAK52083	CLICSB	222	5E-58
	AAD38446	H1 chloride channel; p64H1; CLIC4	221	7E-58
	AAD26136	intracellular chloride channel p64H1	219	3E-57
	AAN76730	chloride channel form B	217	2E-56
	CAC36880	bA802N13.1.1 (chloride intracellular channel 5, isoform 1)	215	5E-56
	BAA91794	unnamed protein product	195	7E-50
NM_025939 Mm.182931 F:2.05 (7to11)	1	$\underline{NP}\ \underline{006443.1}\ \underline{phosphoribosylaninoimidazole}\ \mathrm{carboxylase, phosphoribosylaminoimidazole}\ \mathrm{succinocarboxamide}$	839	0
NP_080215.1		synthetase; phosphoribosylaminoimidazole carboxylase,		
		phosphothosylaminothosylaminothingazore succinocaldoxamine symmerase	3	1
	XP_094999.1	094999.1 similar to Multifunctional protein ADE2	310	4E-84
	XP_116650.2	116650.2 similar to Multifunctional protein ADE2	305	2E-82
J 0 4 6 9 4 Mm.738 F:2.05 (5to11)	1) · 1402236A	collagen alpha1(IV)	563	1E-159
AAA50292.1				
	NP_001836.	_001836.1 alpha 1 type IV collagen preproprotein; collagen IV, alpha-1 polypeptide; collagen of basement	563	1E-159
		membrane, alpha-1 chain		

•		232		
CAC	313153.1 E	CAC13153.1 bA472K17.2 (collagen type IV alpha 1)	563	1E-159
CAA	CAA68698.1 a	alpal-chain	520	1E-146
AA£	A52042.1	AAA52042.1 procollagen alpha-1 type IV	479	1E-133
AAA	A52006.1	pro-alpha-1(IV)	479	1E-133
qpd	pdb 1L11	Chain A, The 1.9-A Crystal Structure Of The Noncollagenous (Nc1) Domain Of Human Placenta Collagen Iv Shows Stabilization Via A Novel Type Of Covalent Met-Lys Cross-Link	474	1E-132
AAD	M97359.1 arresten	arresten	470	1E-130
CAI	B90289.1	dA24A23.1 (collagen, type IV, alpha 5 (Alport syndrome))	422	1E-116
AA	AAA52045.1	collagen type IV alpha 5 chain	422	1E-116
AA	A99480.1	AAA99480.1 alpha-5 type IV collagen	422	1E-116
dN .	203699.1	NP_203699.1 alpha 5 type IV collagen, isoform 2, precursor; collagen IV, alpha-5 polypeptide; collagen of	422	1E-116
dN.	203700.1	NP_203700.1 alpha 5 type IV collagen, isoform 3, precursor; collagen IV, alpha-5 polypeptide; collagen of hasement membrane alpha-5 chain	422	1E-116
NP	000486.1	NP_000486.1 alpha 5 type IV collagen, isoform 1, precursor; collagen IV, alpha-5 polypeptide; collagen of	422	1E-116
AĀ	A51558.1	AAAS1558.1 alpha-5 type IV collagen	422	1E-116
AA	AS1556.1	AAA51556.1 alpha-3 type IV collagen	362	3E-98
AA	A21610.1	AAA21610.1 alpha-3 type IV collagen	362	3E-98
AA	AAF72632.1	tunustatin	362	
ďΝ	000082.1	NP_000082.1 alpha 3 type IV collagen, isoform 1, precursor; collagen IV, alpha-3 polypeptide (goodpasture	362	3E-98

		Courne	hallacen aluka 2/IV) aham meenwear Jang enlice form	362	3E-98
		CUHU3B	conagen aipna 3(1V) cham precursor, rong spince ronn	707	
		CAC36101.1	a3 type IV collagen	362	3E-98
		CAA29075.1	alpha-1 chain precursor (AA -27 to 917) (2953 is 2nd base in codon)	357	2E-96
		AAB19637.1	AAB19637.1 type IV collagen alpha 3 chain	349	3E-94
		pdb 1L11	Chain A, The 1.9-A Crystal Structure Of The Noncollagenous (Nc1) Domain Of Human Placenta	322	4E-86
			Collagen Iv Shows Stabilization Via A Novel Type Of Covalent Met-Lys Cross-Link	5	100
		P08572	Collagen alpha 2(IV) chain precursor	322	4E-86
		NP_001837.1	001837.1 alpha 2 type IV collagen preproprotein; canstatin	322	4E-86
	·	CAA29098.1	alpha (2) chain	322	4E-86
		AAA52043.1	alpha-2 type IV collagen	321	8E-86
		AAA58422.1	collagen alpha-2 type IV	320	1E-85
		AAF72631.1	canstatin	320	1E-85
		CAA20120.1	A20120.1 COLAA6 (Collagen Alpha 6(IV))	307	1E-81
		AAB19038.1	collagen type IV a6 chain	307	1E-81
.,		NP_001838.1	001838.1 type IV alpha 6 collagen, isoform A precursor; collagen IV, alpha-6 polypeptide; collagen of basement membrane, alpha-6	307	1E-81
		Q14031	Collagen alpha 6(IV) chain precursor	307	1E-81
		NP_378667.	378667.1 type IV alpha 6 collagen, isoform B precursor; collagen IV, alpha-6 polypeptide; collagen of basement membrane, alpha-6	307	1E-81
		AAB19039.1	AAB19039.1 collagen type IV a6 chain	307	1E-81

4	
ന	
$^{\circ}$	

1E-170		1E-155	1E-150	1E-132	1E-132	1E-132	1E-131	1E-131	1E-131	1E-130	1E-126	1E-126	1E-113		3E-70	8E-69	1E-68	2E-68	3E-68
603		554	375	330	330	330	327	327	327	327	312	312	414	252	167	166	166	163	164
NP_036250.1 CCR4 carbon catabolite repression 4-like (S. cerevisiae); CCR4-like (carbon catabolite repression	4, S. cerevisiae)	nocturnin	pol protein	polymerase	Gag-Pro-Pol protein	polymerase	pol/eny ORF (bases 3878-8257) first start codon at 4172; Xxx; putative	polymerase	Endogenous retrovirus HERV-K10 putative pol polyprotein [Includes: Reverse transcriptase; Endonuclease]	AAD51793.1 Gag-Pro-Pol-Env protein	AAD51796.1 Gag-Pro-Pol protein	pol protein	nocturnin	gag-pro-pol precursor	polymerase	polymerase	polymerase	polymerase	polymerase
NP_036250.1		AAG01389.1	AAM81188.1 pol protein	AAK11553.1	AAD51797.1	AAD21097.1	AAA88033.1	AAK11554.1	P10266	AAD51793.	AAD51796.	AAL60056.,	AAG01388.,	AAG18012.	AAC63294.1	AAC63291.1	AAC63292.1	AAC63293.1	AAC63290.1
F:2.05 (5to7))																
Mm.86541																	·		,
U70139	AAB62717.1																		

NM_008956 Mm.19117 NP_032982.1		F:2.05 (5to19)	NP_114368.1	114368.1 polypyrimidine tract binding protein, isoform c; RNA binding protein; heterogeneous nuclear ribonucleoprotein polypeptide I; polypyrimidine tract binding protein (heterogeneous nuclear ribonucleoprotein I)	840	0
		,	NP_114367.1	114367.1 polypyrimidine tract binding protein, isoform b; RNA binding protein; heterogeneous nuclear ribonucleoprotein polypeptide I; polypyrimidine tract binding protein (heterogeneous nuclear ribonucleoprotein I)	828	0
			NP_002810.1	002810.1 polypyrimidine tract binding protein, isoform a; RNA binding protein; heterogeneous nuclear ribonucleoprotein polypeptide I; polypyrimidine tract binding protein (heterogeneous nuclear ribonucleoprotein I)	826	0
			BAB71742.1	PTB-like protein L	674	0
			AAM94624.1	AAM94624.1 non-neuronal splice variant nPTB3	019	0
		/	NP_067013.1	polypyrimidine tract binding protein 2; neural polypyrimidine tract binding protein; PTB-like protein	999	0
			AAM94625.1	AAM94625.1 non-neuronal splice variant nPTB4	664	0
			NP_005147.1	005147.1 ROD1 regulator of differentiation 1; fission yeast differentiation regulator; regulator of differentiation (in S. pombi) 1; regulator of differentiation (in S. pombe) 1	640	0
			BAB71743.1	PTB-like protein S	426	1E-118
			XP_063346.3	similar to polypyrimidine-tract binding protein	410	1E-114
			pdb 1QM9	Chain A, Nmr, Representative Structure	312	2E-84
NM_011919 Mm.25709 NP 036049.1	1.25709	F:2.05 (5to19)	AAF07921.1 p33ING1b	p33ING1b	435	1E-122

	1 000000	CM with the contract of the co	434	1F-122
	AAG02578.1	AAGU22/8.1 grown minotroly protein man		
	AAC00501.1	candidate tumor suppressor p33ING1	431	1E-121
	AAG02579.1	growth inhibitory protein ING1	347	9E-96
	NP_005528.2	NP_005528.2 inhibitor of growth family, member 1; inhibitor of growth 1; inhibitor of growth 1 family, member	347	9E-96
	AAF37423.1	ING1 tumor suppressor, variant C	343	2E-94
	AAB60879.1	p33ING1	343	2E-94
	BAA82887.1	p47	343	2E-94
	BAA82889.1	p33	341	7E-94
	BAB08103.1	p24ING1c	309	4E-84
	BAA83462.1	p24 is an alternatively spliced transcript of p33/ING1.	305	7E-83
	CAC20567.1	p32 protein	281	8E-76
	NP_001555.1	_001555.1 inhibitor of growth 1-like	281	1E-75
NM_019447 Mm.27369 F:2.05 (5to19) NP 062320.1	ĀZ	001519.1 HGF activator	1048	0
	CAA93803.1	hepatocyte growth factor (HGF) precursor	1041	0
	NP_000496.1	_000496.1 coagulation factor XII precursor; Hageman factor	462	1E-129
	AAA51986.1	coagulation factor XII	462	1E-129
	AAA70225.1	coagulation factor XII precursor	462	1E-129

ator ssue type isoform I preproprotein; plasminogen activator, tissue type; alteplase; reteplase ama hyaluronic acid binding protein sisue type isoform 3 precursor; plasminogen activator, tissue type; t- sisue type isoform 3 precursor; plasminogen activator, tissue type; t- sisue type isoform 3 precursor; plasminogen activator, tissue type; t- sisue type isoform 3 precursor; plasminogen activator, tissue type; t- sisue type isoform 3 precursor; plasminogen activator, tissue type; t- sisue type isoform 4 precursor; plasminogen activator, tissue type; t- sisue type isoform 4 precursor; plasminogen activator, tissue type; t- sisue type isoform 4 precursor; plasminogen activator, tissue type; t- sisue type isoform 5 precursor; plasminogen activator, tissue type; t- sisue type isoform 5 precursor; plasminogen activator, tissue type; t- sisue type isoform 6 precursor; plasminogen activator, tissue type; t- sisue type isoform a sisue type isoform a sisue type isoform a sisue type; t- sisue type;				237		
BAA00881.1 (issue plasminogen activator, tissue type isoform I preproprotein; plasminogen activator, tissue type; 305				1	370	1E-102
NP_000921.1 plasminogen activator, sisue type isoform 1 preproprotein; plasminogen activator, sisue type; 303 AAB39310.1 plasminogen activator, alteplase, reteplase AAB39310.1 plasminogen activator, alteplase, reteplase NP_004123.1 hyalmonan binding protein 2; hyalmonan binding protein 2; hepatocyte growth factor 300 activator-like protein; plasminogen activator, sisue type isoform 3 precursor, plasminogen activator, sisue type isoform 3 precursor, plasminogen activator, sisue type isoform 3 precursor, plasminogen activator, sisue type isoform 3 precursor, plasminogen activator, sisue type isoform 3 precursor, plasminogen activator, sisue type isoform 3 precursor, plasminogen activator, sisue type isoform 3 precursor, plasminogen activator, sisue type isoform 3 precursor, plasminogen activator, silventation response protein; actual deprivation response; plasminogen activator, silventation response protein; actual deprivation response; plasminogen activator, silventation response protein; actual deprivation response; plasminogen activator, silventation receptor [N-Lough peptide receptor (Romyl peptide receptor) [N-Lough peptide receptor] AAA58481 PMLP-related receptor (M-LP receptor) N-Lough peptide receptor (M-LP receptor) N-Lough peptide receptor (M-LP receptor) N-Lough peptide receptor (M-LP Ross N-Lough Peptide receptor) N-Lough N-Loug			A00881.1	tissue plasminogen activator	305	1E-82
AAB39510.1 plastninogen activator AAB39510.1 plastninogen activator AAB39510.1 plastninogen activator AAB39510.1 plastninogen activator AAB39510.1 plastninogen activator plastninogen			NP_000921.1	plasminogen activator, tissue type isoform 1 preproprotein; plasminogen activator, tissue type; t-plasminogen activator; alteplase; reteplase	305	1E-82
NP_004123.1 hyaluronan binding protein 2; hyaluronan binding protein 2; hepatocyte growth factor 300 activator-like protein; plasma lyaluronan binding protein 2; hepatocyte growth factor 10 activator-like protein; plasma lyaluronan binding protein 10 NP_127509.1 plasminogen activator; tissue type isoform 3 precursor; plasminogen activator; tissue type isoform 3 precursor; plasminogen activator; alteplase. NULL F.2.05 (5to.19) NP_004648.1 serum deprivation response protein; serum deprivation response; phosphatidylserine-binding 611 protein protein protein protein protein protein protein protein protein AAGS48481 FMLP-related receptor-like 1; lipoxin A4 receptor (formyl peptide receptor 11 NP_002021 formyl peptide receptor-like 2 though 1				plasminogen activator	303	6E-82
NULL F.2.05 (5to 19) NP_004648.1 plasminogen activator, itissue type isoform 3 precursor, plasminogen activator, itissue type; to plasminogen activator, alteplase, reteplase Protein			NP_004123.1	hyaluronan binding protein 2; hyaluronic acid binding protein 2; hepatocyte growth factor activator-like protein: plasma hyaluronan binding protein	l.,	7E-81
NULL F.2.05 (5to19) NP_004648.1 serum deprivation response protein; serum deprivation response; phosphatidylserine-binding 611			NP_127509.1	plasminogen activator, tissue type isoform 3 precursor; plasminogen activator, tissue type; t-		2E-80
NULL F:2.05 (5to 19) NP_004648.1 serum deprivation response protein; serum deprivation response; phosphatidylserine-binding 611				piasimiogen achvator, arcepiase, recepiase		
AAG27093.1 feucine-zipper protein FKSG13 215 F.2.04 (YioO) NP_001453 formyl peptide receptor-like 1; lipoxin A4 receptor (formyl peptide receptor II 501 AAA52474 DEFINITION N-formyl peptide receptor-like 2 protein 419 NP_002021 formyl peptide receptor-like 2 410 NP_002020 formyl peptide receptor (fMLP receptor) (N-formyl peptide receptor) NP_002020 formyl peptide receptor (fMLP receptor) (N-formyl peptide receptor) AAA36362 N-formyl peptide receptor (fMLP-R98 407 AAA36362 N-formyl peptide receptor fMLP-R98 408 AAA36362 N-formyl peptide receptor fMLP-R98 409 AAA36362 N-formyl peptide receptor fMLP-R98 409 AAA36362 N-formyl peptide receptor fMLP-R98 409 AAA36363 Chemokine-like receptor fMLP-R98 400 AA		F:2.05 (5to19		serum deprivation response protein; serum deprivation response; phosphatidylserine-binding protein	·	1E-174
F.2.04 (Y10C) NP_001453 formyl peptide receptor-like I; lipoxin A4 receptor (formyl peptide receptor related) 502			AAG27093.1	leucine-zipper protein FKSG13	215	3E-55
AAA58481 FMLP-related receptor II AAA52474 DEFINITION N-formyl peptide receptor-like 2 protein NP 002021 formyl peptide receptor 1 NP 002020 formyl peptide receptor 1 P21462 fMet-Leu-Phe receptor (fMLP receptor) (N-formyl peptide receptor) AA2009 N-formyl peptide receptor AAA36362 N-formyl peptide receptor fMLP-R98 AAA36362 N-formylpeptide receptor fMLP-R98 AAA36363 chemokine-like receptor 1	008039 Mm.571		1 1	formyl peptide receptor-like 1; lipoxin A4 receptor (formyl peptide receptor related)	502	1E-142
formyl peptide receptor-like 2 protein formyl peptide receptor-like 2 formyl peptide receptor 1 formyl peptide receptor 1 formyl peptide receptor 1 formyl peptide receptor (fMLP receptor) (N-formyl peptide receptor) Chemoattractant receptor). N-formyl peptide receptor N-formyl peptide receptor FPR) (N-formylpeptide receptor) N-formylpeptide receptor fMLP-R98 orphan G-protein coupled receptor Dez isoform a chemokine-like receptor 1	32065.1		A A A 50401	DMT D related tonewher II	501	1F-142
formyl peptide receptor-like 2 formyl peptide receptor 1 formyl peptide receptor (fMLP receptor) (N-formyl peptide receptor) (FPR) (N-formylpeptide chemoattractant receptor). N-formyl peptide receptor N-formylpeptide receptor fMLP-R98 orphan G-protein coupled receptor Dez isoform a chemokine-like receptor 1			AAA52474	DEFINITION N-formyl peptide receptor-like 2 protein	419	
formyl peptide receptor 1 fMet-Leu-Phe receptor (fMLP receptor) (N-formyl peptide receptor) (FPR) (N-formylpeptide chemoattractant receptor). N-formyl peptide receptor N-formylpeptide receptor fMLP-R98 orphan G-protein coupled receptor Dez isoform a chemokine-like receptor 1			NP 002021	formyl peptide receptor-like 2	415	1E-116
fMet-Leu-Phe receptor (fMLP receptor) (N-formyl peptide receptor) (FPR) (N-formylpeptide chemoattractant receptor). N-formyl peptide receptor N-formylpeptide receptor fMLP-R98 orphan G-protein coupled receptor Dez isoform a chemokine-like receptor 1			NP 002020	formyl peptide receptor 1	410	1E-114
chemoattractant receptor). N-formyl peptide receptor N-formylpeptide receptor fMLP-R98 orphan G-protein coupled receptor Dez isoform a chemokine-like receptor 1			P21462	fMet-Leu-Phe receptor (fMLP receptor) (N-formyl peptide receptor) (FPR) (N-formylpeptide		1E-113
N-formyl peptide receptor N-formylpeptide receptor fMLP-R98 orphan G-protein coupled receptor Dez isoform a chemokine-like receptor 1				chemoattractant receptor).		
N-formylpeptide receptor fMLP-R98 orphan G-protein coupled receptor Dez isoform a chemokine-like receptor 1			A42009	N-formyl peptide receptor	9	1E-113
N-formylpeptide receptor fMLP-R98 orphan G-protein coupled receptor Dez isoform a chemokine-like receptor 1					Apı	
orphan G-protein coupled receptor Dez isoform a chemokine-like receptor 1			AAA36362		404	
chemokine-like receptor 1			AAC51258		201	
			NP 004063		201	2E-51

Ŋ

		099788	2.38 Chemokine receptor-like 1 (G-protein coupled receptor DEZ) (G protein-coupled receptor	201	2E-51
			ChemR23).		
NM_009417 Mm.4991	F:2.04 (YtoM)	P07202	Thyroid peroxidase precursor (TPO).	1414	0
NP 033443.1					·
		AAA61215	thyroid peroxidase	1412	0
		CAA68467	precursor polypeptide	1409	0
		NP 000538	thyroid peroxidase isoform a; thyroperoxidase; thyroid microsomal antigen	1409	0
		AAA61217	thyroid peroxidase	1404	0
		NP 783651	thyroid peroxidase isoform c; thyroperoxidase; thyroid microsomal antigen	1377	0
		NP 783652	thyroid peroxidase isoform d; thyroperoxidase; thyroid microsomal antigen	1311	0
		AAA61216	thyroid peroxidase	1251	0
		NP 783650	thyroid peroxidase isoform b; thyroperoxidase; thyroid microsomal antigen	1248	0
		CAA35235	thyroid peroxidase (AA 1-876)	1246	0
		AAL74416	_	741	
		NP 783653		738	0
		CAA33438	_	593	1E-169
		NP 000241	•	593	1E-169
	-	C28894	$\overline{}$	583	1E-166
		NP 000493	_	582	1E-166
		B28894	1	558	1E-158
		CAA32530	_	550	
		BAA13219		536	1E-152
		XP 056455		536	
		XP 042207	similar to Lactoperoxidase precursor (LPO) (Salivary peroxidase) (SPO)	530	1E-150
		AAN04473		505	1E-142
		ICXPC	-	454	
		IMYPC	1	454	
		AAN04474	thyroid peroxidase isoform 2/4	437	1E-122
		AAN04471		398	1E-110
		AAA61218	thyroperoxidase	368	
		AAA61219	lhyroperoxidase	336	
•		AAA63213.	lactoperoxidase	321	1.4E-86
					•

Ŋ

				239		
			NP 653252	25471	296	296 1.9E-79
M_007472 Mm.18625	Im.18625	F:2.04 (7to11)	152366	uterine water channel	496	1E-140
r 031498.1			AAH22486.1	aquaporin 1 (channel-forming integral protein, 28kD)	495	1E-139
			NP_000376.1	000376.1 aquaporin 1; aquaporin 1 (channel-forming integral protein, 28kDa); Aquaporin-1 (channel-	495	1E-139
				forming integral protein, 28kDa); Colton blood group		
			AAL87136.1		488	1E-137
			AAC50649.1	channel-like integral membrane protein	293	7E-79
			AAC23788.1	aquaporin	276	8E-74
			AAC03168.1	putative alternative lens membrane intrinsic protein	238	2E-62
			NP_036196.1	_036196.1 major intrinsic protein of lens fiber; aquaporin	233	1E-60
			NP_000477.1	_000477.1 aquaporin 2; Aquaporin-2 (collecting duct)	230	9E-60
			AAB30268.1	hAQP-CD=collecting duct aquaporin [human, kidney, Peptide, 271 aa]	228	3E-59
			151877	water-channel aquaporin 2	227	7E-59
			164818	water-channel aquaporin 2	227	7E-59
			AAC16481.1	aquaporin (water channel protein)	223	1E-57
			NP_004019.1	aquaporin 4 C2 isoform; mercurial-insensitive water channel	221	3E-57
			NP_001641.1	l aquaporin 4 isoform a; mercurial-insensitive water channel	221	3E-57
			139177	mercurial-insensitive water channel	221	4E-57
			139178	aquaporin 4, long splice form	221	4E-57
			NP_001642.1	001642.1 aquaporin 5; Aquaporin-5	218	5E-56

15 10 10

\smile	
d,	
S	

			047	503	170
NM_029239 Mm.108699 F:2.04 (7to11)	F:2.04 (7to11)	NP_005804.1	005804.1 protein kinase C, nu; serine-threonine protein kinase	594	0/1-31
NP 083515.1					, ,
		AAH30706.1	Similar to protein kinase C, nu	592	1E-169
		NP_002733.1	protein kinase C, mu	378	1E-105
		NP_057541.2	057541.2 protein kinase D2	342	6E-94
		BAC11508.1	unnamed protein product	218	2E-56
)		BAC11127.1	unnamed protein product	218	2E-56
3 0 Mm.279	F:2.04 (7to19)	NP_057146.1	057146.1 CGI-128 protein *	298	4E-80
BAB23024.1					
					77. 44.
NM_020520 Mm.29666	F:2.04 (7to19)	CAB55356.1	carnitine/acylcarnitine translocase	582	1E-166
NP 065266.1				- 1	77. 4.
		NP_000378.1	NP_000378.1 carmitine/acylcarmitine translocase; Carmitine-acylcarmitine translocase; carmitine-acylcarmitine	281	11-1100
			carrier; solute carrier family 25 (camitine/acylcamitine translocase), member 20		
A K 0 0 7 2 6 4 Mm.200370 F:2.04 (5to19)	F:2.04 (5to19)	AAD12227.1	AAD12227.1 similar to uridine phosphorylase, similar to Q16831 (PID:g2494059)	447	1E-125
BAB24924.1					
		XP_087230.2	087230.2 similar to Uridine phosphorylase (UDRPase)	428	1E-120
		NP_003355.1	uridine phosphorylase	316	3E-86
A K 0 0 8 0 9 8 Mm.10706	F:2.04 (5to19)	NP_116024.1	_116024.1 seven transmembrane domain protein	442	1E-123
1,000,000		CAA77013.1	seven transmembrane domain protein	414	1E-115
				_	

Ŋ

10

_	7
$\overline{}$	r
C	٧

		741		
NM_011017 Mm.200907 F:2.04 (5to19)	AAG17977.1	ипклоwп	561	1E-159
NP 035147.1	·			
	NP_055067.1	055067.1 ornithine transporter 1 (hyperornithinemia-hyperammonemia-homocitrullinuria); ornithine	le 560	1E-159
		transporter 1	_	
	NP_114153.1	ornithine transporter 2	499	1E-141
	AAM94902.1	AAM94902.1 ornithine transporter 2	497	1E-140
NM_029796 Mm.176946 F:2.04 (5to19)	NP_443204.1	443204.1 leucine-rich alpha-2-glycoprotein	330	3E-90
NP 084072.1				
	NBHUA2	leucine-rich alpha-2-glycoprotein	329	6E-90
	AAH34389.1	leucine-rich alpha-2-glycoprotein	327	2E-89
NM_021532 Mm.46662 F.2.03 (YtoO)	NP_057735	DAPPER1; heptacellular carcinoma novel gene 3	970	0
NP 067507.2			_	
	AAF65569	heptacellular carcinoma novel gene-3 protein	714	0
	CAD61905	unnamed protein product	714	0
NM_011087 Mm.193462 F:2.03 (YtoO)	NP_077294	immunoglobulin-like transcript 8	407	1E-113
NP_035217.1			_	
	AAC51892	immunoglobulin-like transcript 5 protein	400	1E:111
	AAC51902	immunoglobulin-like transcript 5	400	1E-111
-	AAC51893	immunoglobulin-like transcript 5 protein	400	1E-111
	AAB88120	immunoglobulin-like transcript 5; ILT5	399	1E-110
	AAB87667	leucocyte immunoglobulin-like receptor-3; LIR-3	399	1E-110
	AAC51888	immunoglobulin-like transcript 5 protein	399	1E-110
	AAC51894	immunoglobulin-like transcript 5 protein	399	1E-110
	AAC51889	immunoglobulin-like transcript 5 protein	399	1E-110

S

0

H

 \sim

\sim	:
ч	•
C	į

		747	000	11. 110
	AAC51895	immunoglobulin-like transcript 5 protein	252	11-11
	AAC51901	immunoglobulin-like transcript 5	397	1E-110
	NP_006855	leukocyte immunoglobulin-like receptor, subfamily B (with TM and ITIM dornains), member 3;	396	1E-109
		leukocyte immunoglobulin-like receptor 3		
	AAC51896	immunoglobulin-like transcript 5 protein	396	1E-109
	AAC51890	immunoglobulin-like transcript 5 protein	395	1E-109
	AAC51891	immunoglobulin-like transcript 5 protein	395	1
	AAC51900	immunoglobulin-like transcript 5	394	1E-109
	AAC51897	immunoglobulin-like transcript 5 protein	394	1E-109
	AAC51887	immunoglobulin-like transcript 5 protein	393	
	NP 036408	immunoglobulin-like transcript 7	393	1E-108
	AAL36993	immunoglobulin-like transcript-7	382	1E-105
	AAC51178	immunoglobulin-like transcript 1c	382	1E-105
	AAD50364	immunoglobulin-like transcript 1c	382	1E-105
	AAD17990	immunoglobulin-like transcript 1c variant 3	380	1E-105
	AAD50365	immunoglobulin-like transcript 1c	380	1E-105
	AAD17991	inmunoglobulin-like transcript 1c variant 4	380	1E-105
	AAC51176	innmunoglobulin-like transcript la	376	1E-103
	JC5897	killer cell inhibitory receptor p91 precursor	376	1E-103
	NP_006854	leukocyte immunoglobulin-like receptor, subfamily A (with TM domain), member 1; leukocyte	375	1E-103
		immunoglobulin-like receptor 6		
	NP_006857	leukocyte immunoglobulin-like receptor, subfamily A (with TM domain), member 2; leukocyte	375	1E-103
		immunoglobulin-like receptor 7	丄	
	NP_006831	leukocyte immunoglobulin-like receptor, subfamily B (with TM and ITIM domains), member 5	371	
	AAM18038	leucocyte immunoglobulin-like receptor	359) 2E-98
	AAM18036	leucocyte immunoglobulin-like receptor	358	3 2E-98
	AAC51885	immunoglobulin-like transcript 6	358	
	NP_006856	leukocyte immunoglobulin-like receptor, subfamily A (without TM domain), member 3;	358	8 4E-98
		leukocyte immunoglobulin-like receptor 4		

(ግ
4
à

		243	157	8B 08
	AAM18040	leucocyte immunogiobulin-like receptor	100	02-70
	AAM18041	leucocyte immunoglobulin-like receptor	357	8E-98
	AAM18035	leucocyte immunoglobulin-like receptor	356	1E-97
	AAM18037.	leucocyte immunoglobulin-like receptor	356	1E-97
	AAH28208	leukocyte immunoglobulin-like receptor, subfamily A (without TM domain), member 3	356	1E-97
	AAB87661	leucocyte inmunoglobulin-like receptor-4; LIR-4	355	3E-97
	AAB68667	monocyte inhibitory receptor precursor	353	7E-97
	AAH36827	Unknown (protein for MGC:46153)	352	2E-96
	NP_005865	leukocyte immunoglobulin-like receptor, subfamily B (with TM and ITIM domains), member 2;	352	2E-96
		leukocyte immunoglobulin-like receptor 2		
	AAL36990	leukocyte immunoglobulin-like receptor-2	352	2E-96
	AAC51883	immunoglobulin-like transcript 4	352	2E-96
	AAC51880	inmunoglobulin-like transcript 2b	351	4E-96
	AAL36991	leukocyte immunoglobulin-like receptor-2	350	6E-96
	AAB88119	immunoglobulin-like transcript 4; ILT4	350	8E-96
	AAB67711	MIR-10	350	8E-96
	NP_006660	leukocyte immunoglobulin-like receptor, subfamily B (with TM and ITIM domains), member 1;	349	1E-95
-		leukocyte immunoglobulin-like receptor 1; CD85 antigen		
	AAC51879	immunoglobulin-like transcript 2a	345	2E-94
	AAG08984	leukocyte immunoglobulin-like receptor 1	345	2E-94
	AAB63522	leucocyte immunoglobulin-like receptor-1	345	2E-94
	AAC51881	immunoglobulin-like transcript 2c	345	2E-94
	AAL36989	leukocyte immunoglobulin-like receptor-1	345	2E-94
	AAB67710	MIR-7	345	2E-94
	AAL36988	leukocyte immunoglobulin-like receptor-1	345	2E-94
	XP_115639	similar to immunoglobulin-like transcript 8	271	6E-72
	NP_077293	leukocyte immunoglobulin-like receptor, subfamily A, member 5; immunoglobulin-like transcript	t 260	8E-69
	A A C 007.62	10 immunoclopulin like transcript 10 protein	258	SB-68
	70/66000	Infilmatiogrammatic transcript to protein	222	-

.

0

7

 $\overset{\circ}{\sim}$

`	_		244	ľ	
		BAC03380	FLJ00275 protein	241	4E-63
		BAB71361	unnamed protein product	241	6E-63
		AAC51886	immunoglobulin-like transcript 6a	218	6E-56
NM_019922 Mm,20904	F:2.03 (11to19)	AAH08745.1	F.2.03 (11to19) AAH08745.1 cartilage associated protein	999	0
NP 064306.1					
		NP_006362.1	NP_006362.1 cartilage associated protein; cartilage-associated protein	999	0
	;				
		BAC03743.1	unnamed protein product	633	1E-180
		CAC16786.1	nucleolar protein No55	403	1E-111
		NP_006446.1	NP_006446.1 nucleolar autoantigen (55kD) similar to rat synaptonemal complex	402	1E-111
A F 3 8 5 6 8 2 Mm.27242 AAK63363 1	F:2.03 (7to11)	NP_071442.1	NP_071442.1 EGF-TM7-latrophilin-related protein	934	0
		BAA34488.1	KIAA0768 protein	359	2E-98
		NP_056051.1 lectomedin-3	lectomedin-3	348	4E-95
		AAD54676.1	lectomedin-1 beta	341	4E-93
		NP_036434.1	latrophilin 1; KIAA0786 protein; lectomedin-1; latrophilin	341	4E-93
		AAD54675.1	AAD54675.1 lectomedin-1 alpha	341	4E-93
		BAA34506.1	BAA34506.1 KLAA0786 protein	337	8E-92
		AAG27461.1 lectomedin-2	lectomcdin-2	330	1E-89
		NP_055736.1	lectomedin-2; KIAA0821 protein	330	1E-89
		AAH07587.1	AAH07587.1 Unknown (protein for IMAGE:3162852)	322	2E-87

S

0

1.5

 \approx

2.4

				245		
			NP_690880.1	P_690880.1 egf-like module containing, mucin-like, hormone receptor-like sequence 2 isoform b	281	4E-75
			NP_001775.2	P_001775.2 CD97 antigen, isoform 2 precursor; leukocyte antigen CD97; seven-span transnembrane protein	280	2E-74
			NP_115960.1	NP_115960.1 egf-like module-containing mucin-like receptor 3 isoform a	278	5E-74
			AAF21974.1	EGF-like module EMR2	277	8E-74
			NP_038475.2	NP_038475.2 legf-like module containing, mucin-like, hormone receptor-like sequence 2 isoform a	275	4E-73
			BAC06146.1	seven transmembrane helix receptor	275	4E-73
			137225	leucocyte antigen CD97	275	5E-73
			NP_510966.1	CD97 antigen, isoform 1 precursor; leukocyte antigen CD97; seven-span transmembrane protein	265	3E-70
			AAB36682.1	CD97	265	3E-70
			BAC06178.1	seven transmembrane helix receptor	265	3E-70
			BAC06133.1	seven transmembrane helix receptor	260	1E-68
			P48960	Leucocyte antigen CD97 precursor	260	1E-68
			NP_001965.1	egf-like module containing, mucin-like, hormone receptor-like sequence 1; egf-like module	259	2E-68
				containing, mucin-like, hormone receptor-like		
			NP_690881.1	egf-like module containing, mucin-like, hormone receptor-like sequence 2 isoform c	254	7E-67
:			NP_690883.1	egf-like module containing, mucin-like, hormone receptor-like sequence 2 isoform e	231	6E-60
			NP_690882.1	NP 690882.1 egf-like module containing, mucin-like, hormone receptor-like sequence 2 isoform d	231	6E-60
			NP_690885.1	egf-like module containing, mucin-like, hormone receptor-like sequence 2 isoform g	231	6E-60
			NP_690884.1	egf-like module containing, mucin-like, hormone receptor-like sequence 2 isoform f	231	6E-60
			AAC05172.1 R29368_2	R29368_2	225	4E-58
NM_008625 Mm.2019	lm.2019	F:2.03 (7to 19)	NP_002429.1	NP_002429.1 mannose receptor C type 1 precursor; mannose receptor precursor; macrophage mannose receptor	2521	0
NP_032651.1				-		
			NP_006030.1	P_006030.1 mannose receptor, C type 2; KIAA0709 gene product; endocytic receptor (macrophage mannose	831	0
				receptor family), likely ortholog of mouse mannose receptor, C type 2		

5

7

7

				246	,	(
			AAD30280.1	AAD30280.1 endocytic receptor Endo180	830	5 –
			NP_031392.2	031392.2 phospholipase A2 receptor 1, 180kDa; phospholipase A2 receptor 1, 180kD	573	1E-162
r coood year	N. 1510	T. 7 02 (71, 10)	MD 000040 1	000040 1 ATD Linding cassette sub-femily D member 3. Perovisoms membrane protein-[(70k1)]:	1214	0
WIN _ 000991 MIN _ 1	MIII. 1319	(7.017) (7.017)	11. CF0200_1VI			
NP 033017.1				peroxisomal membrane protein 1 (70kD, Zellweger syndrome); peroxisomal membrane protein-1	!	1
			S20313	peroxisonal membrane protein, 70K	1207	0
			CAA58470.1	70kD peroxisomal integral membrane protein	1155	0
)24.2	ATP-binding cassette, sub-family D (ALD), member 1; adrenoleukodystrophy protein	445	1E-124
			JC5712	adrenoleukodystrophy related proteiń	444	1E-124
			A	adrenoleukodystrophy protein	443	1E-124
			P33897	Adrenoleukodystrophy protein (ALDP)	443	1E-124
			NP_005155.1	ATP-binding cassette, sub-family D, member 2; adrenoleukodystrophy-like 1; hALDR	442	1E-124
NM_025422 Mm.30109	Mm.30109	F:2.03 (7to19)	NP_055695.1	NP_055695.1 KIAA0022 gene product	251	2E-66
NP 079698.1						
NM_007624 Mm.28148	Mm.28148	F:2.03 (5to19)	NP_009207.2	NP_009207.2 chromobox homolog 3; heterochromatin protein HP1 gamma; HP1 gamma homolog;	261	2E-69
NP 031650.1				heterochromatin-like protein 1		
			AAF62370.1	heterochromatin-like protein l	261	3E-69
			Q13185	Chromobox protein homolog 3 (Heterochromatin protein 1 homolog gamna) (HP1 gamma)	259	1E-68
				(Modifier 2 protein) (HECH)		
			AAB48101.1	HP1Hs-gamma	259	1E-68
			NP_006798.1	chromobox homolog 1 (HP1 beta homolog Drosophila); heterochromatin protein p25 beta;	; 199	1E-50
				chromobox homolog 1 (Drosophila HP1 beta)		
NM_013762 Mm.3486	Mm.3486	F:2.03 (5to 19)	NP_000958.1	NP_000958.1 ribosomal protein L3; 60S ribosomal protein L3; HIV-1 TAR RNA-binding protein B	783	0
NP 038790.1						

マ	
S	

			1.42		
		AAH08492.1	AAH08492.1 ribosomal protein L3	780	
		184501	ribosomal protein L3 - human (fragment).	773	0
		22790.1	Unknown (protein for IMAGE:3538792)	724	0
		CAB76201.1	ribosomal protein L3	662	0
		NP 005052.1	ribosomal protein L3-like; 60S ribosomal protein L3-like	989	0
		AAK61301.1	60S ribosomal protein L3 like	634	0
		XP 172501.1	172501.1 similar to ribosomal protein L3; 60S ribosomal protein L3; HIV-1 TAR RNA-binding protein B	625	1E-179
		AAA91344.1	ARBP-b gene product	532	1E-151
		AAH04323.1	Similar to RIKEN cDNA 1110057H16 gene	448	1E-126
		XP_070263.1	similar to ribosomal protein L3; 60S ribosomal protein L3; HIV-1 TAR RNA-binding protein B	419	1E-117
NM_013837 Mm.16084	F:2.03 (5to19)	NP_003587.1	003587.1 tyrosylprotein sulfotransferase 1.	734	0
NP 038865.1		NP 003586.1	tyrosylprotein sulfotransferase 2; Tyrosylprotein phosphotransferase 2	486	1E-137
		CAB66558.1	hypothetical protein	481	1E-135
NM_016751 Mm.3115 NP_0580311	F:2.03 (5to19)	XP_092649.1	092649.1 similar to Kupffer cell receptor	209	1E-144
		BAC04786.1	unnamed protein product	433	1E-121
		NP_056532.1	056532.1 Langerhans cell specific c-type lectin; langerin	168	3E-41
NM_008043 Mm.4573 NP 032069.1	F:2.02 (YtoM)	NP_005470.	frequently rearranged in advanced T-cell lymphomas	312	2E-84

10 10 15

		ſ	0.47	210	07: 0.4
			GSK-3beta binding protein FRAT1	OIC.	0.5-04
		AAH34476	frequently rearranged in advanced T-cell lymphomas	308	3E-83
NM_008905 Mm.2817 F;2.	F;2.02 (7to19)	CAD39087.1	hypothetical protein	1462	0
NP 032931.1		-			
		AAH21714.1	imilar to protein tyrosine phosphatase, receptor-type, F interacting protein, binding protein 2	1455	0
		XP_084578.4	XP_084578.4 similar to hypothetical protein	1453	0
		AAC26104	liprin-beta2	1299	0
		NP_003613.1	PTPRF interacting protein, binding protein 1 (liprin beta 1)	742	0
		BAA86544.2	BAA86544.2 KIAA1230 protein	663	0
NM_030693 Mm.1566 F:2. NP 109618.1	F:2.02 (7to19)	AAH05174.1	activating transcription factor 5	177	5E-44
		NP_036200.2	036200.2 activating transcription factor 5	164	3E-40
NM_008280 Mm.362 F:2 NP 032306.1	F:2.02 (5to 19)	NP_000227.1	_000227.1 lipase C preciusor	780	0
		AAA59520.1	hepatic lipase precursor	778	0
		A28997	triacylglycerol lipase (EC 3.1.1.3) precursor, hepatic	777	0
		NP_006024.1	_006024.1 endothelial lipase precursor; endothelial cell-derived lipase	415	1E-114
		NP_000228.1	NP_000228.1 lipoprotein lipase precursor	396	1E-109
	-	AAH11353.1	Similar to lipoprotein lipase	395	1E-109
		AAC61679.1	lipoprotein lipase precursor	311	2E-83

Ŋ

0

H

	7 F-2 02 (Sto 19)	NP 002208 1	NP 002208 Thre-slips (globulin) inhibitor H3 nolyneptide: Inter-slips (globulin) inhibitor H3 nolyneptide 1471	1471	
NP 032433.1		ı			<u> </u>
		S30350	inter-alpha-trypsin inhibitor heavy chain 3 precursor	1471	0
		P19827	Inter-alpha-trypsin inhibitor heavy chain H1 precursor (ITI heavy chain H1) (Inter-alpha-inhibito	911	0
			heavy chain 1) (Inter-alpha-trypsin inhibitor complex component III) (Serum-derived hyaluronan-		
			associated protein) (SHAP)		
		S24391	inter-alpha-trypsin inhibitor heavy chain H1 precursor	806	0
		NP_002206.1	NP_002206.1 inter-alpha (globulin) inhibitor, H1 polypeptide	206	0
		S04484	inter-alpha-trypsin inhibitor chain 3 - human	878	0
		CAA32821.1	lambda HuHITI-13	878	0
		CAA34346.1	CAA34346.1 inter-alpha-trypsin inhibitor C-terminal	872	0
		P19823	Inter-alpha-trypsin inhibitor heavy chain H2 precursor (ITI heavy chain H2) (Inter-alpha-inhibitor	699	0
	· · · · · · · · · · · · · · · · · · ·		$heavychain2)(Inter-alpha-tryps ininhibitorcomplexcomponent\Pi)(Serum-derivedhyaluronan-derived hyaluronan-derived hyal$		
			associated protein) (SHAP)		
		NP_002207.1	NP_002207.1 inter-alpha (globulin) inhibitor, H2 polypeptide	664	0
		Q14624	Inter-alpha-trypsin inhibitor heavy chain H4 precursor (ITI heavy chain H4) (Inter-alpha-inhibitor	602	1E-170
			heavy chain 4) (Inter-alpha-trypsin inhibitor family heavy chain-related protein) (IHRP) (Plasma		
			kallikrein sensitive glycoprotein 120) (PK-120) (GP120) (PRO1851) [Contains: GP57]		
		AAD05198.1	inter-alpha-trypsin inhibitor family heavy chain-related protein	601	1E-170
/		BAA07536.1	BAA07536.1 PK-120 precursor	601	1E-170
		NP_002209.1	NP_002209.1 inter-alpha (globulin) inhibitor H4 (plasma Kallikrein-sensitive glycoprotein); inter-alpha	601	1E-170
			(globulin) inhibitor, H polypeptide-like 1; Inter-alpha (globulin) inhibitor, H4 polypeptide		
NM_009254 Mm.2623	3 F:2.02 (5to19)	NP_004559.2	NP_004559.2 serine (or cysteine) proteinase inhibitor, clade B (ovalbumin), member 6; protease inhibitor 6	549	1E-156
NP 033280.1			(placental thrombin inhibitor		

				548	1E-156
		A48681	placental thrombin inhibitor	L	100
		NP_002631.1	002631.1 ne (or cysteine) proteinase inhibitor, clade B (ovalbumin), member 8; protease inhibitor 8	459	15-179
		NP 004146.1	(ovalbumin type) NP 004146.1 serine (or cysteine) proteinase inhibitor, clade B (ovalbumin), member 9; protease inhibitor 9	445	1E-125
		1	(ovalbumin type)		
		NP_109591.1	109591.1 e (or cysteine) proteinase inhibitor, clade B (ovalbumin), member 1; protease inhibitor 2 (anti-	330	3E-90
			elastase), monocyte/neutrophil; protease inhibitor 2 (anti-clastase), monocyte/neutrophil derived		
		XP 036951.4	036951.4 similar to Squamous cell carcinoma antigen 2 (SCCA-2) (Leupin	327	2E-89
		T38202	leupin precursor - human	327	2E-89
		138201	squamous cell carcinoma antigen 1 - human	325	7E-89
ō		NP_008850.1	008850.1 serine (or cysteine) proteinase inhibitor, clade B (ovalbumin), member 3; squamous cell	1 325	9E-89
			carcinoma antigen 1		
		JT0966	squamous cell carcinoma antigen	325	9E-89
		pdb/1JRR.	Chain A, Human Plasminogen Activator Inhibitor-2.[loop (66-98) Deletionmutant] Complexed	d 317	2E-86
			With Peptide Mimicking The Reactive Center Loop	_	
		AAH34528.1	Similar to serine (or cysteine) proteinase inhibitor, clade B (ovalbumin), member 8	313	3E-85
		NP_002566.1	NP_002566.1 serine (or cysteine) proteinase inhibitor, clade B (ovalbumin), member 2; plasminogen activator	ır 306	6E-83
			inhibitor, type II (arginine-serpin)		
		NP_005015.1		0 304	2E-82
			(ovalbumin type, bomapin)		
		BAB40773.1	SCCA2b	303	3E-82
		AAH12609.1	Similar to serine (or cysteine) proteinase inhibitor, clade B (ovalbumin), member 2	303	5E-82
		AAA36413.1	plasminogen activator inhibitor	301	1E-81
		JC7118	headpin serine proteinase inhibitor	285	8E-77
		CAA04937.1 hurpin	hurpin	285	8E-77
NM 009658 Mm 451	E-2 ()2 (5tn 10)	NP 001619	D 001619 1 aldo-keto reduciase family I, member B1; aldehyde reductase 1; aldose reductase; low Km aldose	se 542	IE-154

11 Wild Type Complexed With Nadp+ And Glucose-6- 540 11 1.21) Mutant With Tyr 48 Replaced By His (Y48h) 538 11 2.21) Mutant With Cys 298 Replaced By Ser (C298s) 536 11 2.21) Mutant With Cys 298 Replaced By Ser (C298s) 536 11 2.22 MUTANT HUMAN ALDOSE Reductase family 448 13 3.23 Bl0; aldose reductase-like peptide; aldose reductase-related aldose reductase-like peptide; aldose reductase-like 337 11 3.32 Bl1 (aldose reductase-like 318 310 3.32 Eto reductase related protein 310 3.32 Eto reductase related protein 310 3.33 Al; aldehyde reductase, alcohol dehydrogenase 297 3.34 Steroid-5-beta-reductase, beta polypeptide 1 (3-oxo-5 293 12 13 13 3.34 Steroid 5-beta-reductase 291 291 3.35 Sata 1); steroid 5-beta-reductase 291 291 3.36 Sata 1); steroid 5-beta-reductase 291 291 291 291 291 291 201	-	LOJ.		
Chain, Aldose Reductase (E.C.1.1.21) Wild Type Complexed With Nadp+ And Glucose-6. 540 11 Phosphate Chain, Aldose Reductase (E.C.1.1.121) Mutant With Tyr 48 Replaced By His (Y48lt) 538 11 Complexed With Nadp+ And Cirate Chain, Aldose Reductase (E.C.1.1.1.21) Mutant With Cys 298 Replaced By Ser (C298s) 536 11 Complex With Nadph Interstine reductase family 1, member B10; aldose reductase-like peptide; aldo-keto reductase family 1, member B11 (aldose reductase-like aldo-keto reductase family 1, member B11 (aldose reductase-like aldo-keto reductase family 1, member B11 (aldose reductase-like aldo-keto reductase family 1, member B11 (aldose reductase related protein Chain, Aldehyde Reductase Chain Aldehyde reductase family 1, member B1; steroid 5-beta-reductase, beta polypeptide 1 (3-0xo-5- Chain Aldehyde re		reductase; Lii>-2 CICL tumor antigen		
Chain , Aldose Reductase (E.C.1.1.21) Wild Type Complexed With Nadp+ And Glucose-6-540 II Phosphate Chain , Aldose Reductase (E.C.1.1.121) Mutant With Tyr 48 Replaced By His (Y48h) 538 II Complexed With Nadp+ And Citrate Complexed With Nadp+ And Citrate Complexed With Nadp+ And Citrate Complexed With Nadp+ Citrate Complex With Nadp+ And Citrate Complex With Nadp+ And Citrate Complex With Nadp+ And Citrate Complex With Nadp+ And Citrate Complex With Nadp+ And Citrate Complex With Nadp+ And Citrate Complex With Nadp+ And Citrate Complex With Nadp+ And Citrate Complex With Nadp+ And Citrate Complex With Nadp+ And Citrate Complex With Nadp+ And Citrate Complex With Nadp+ And Citrate Complex With Nadp+ And Citrate Complex With Nadp+ And Citrate Complex With Nadp+ And Citrate Complex With Nadp+ And Citrate Complex With Nadp+ And Citrate Citrate Algore Reductase 1.1 aldo-keto reductase family 1, member B10, aldose reductase-like peptide; aldose reductase family 1, member B11 (aldose reductase-like peptide) 3.1 similar to aldo-keto reductase family 1, member B11 (aldose reductase aloohol dehydrogenase Chain , Aldehyde Reductase loopADR; aldo-keto reductase related protein Chain , Aldehyde Reductase family 1, member A1; aldehyde reductase, beta polypeptide 1 (3-oxo-5 207 Chain , Aldehyde Reductase family 1, member A1; aldehyde reductase, beta polypeptide 1 (3-oxo-5 208 beta-steroid delta 4-dehydrogenase beta 1); steroid 5-beta-reductase aldehyde reductase	AAA51714.1	aldose reductase	541	1E-154
Chain , Aldose Reductase (E.C.1.1.121) Mutant With Tyr 48 Replaced By His (Y48h) 538 11 Complexed With Nadp+ And Citrate Complexed With Nadp+ And Citrate Chain , Aldose Reductase (E.C.1.1.21) Mutant With Cys 298 Replaced By Ser (C298s) 536 11 Complex With Nadph Complex With Nadph Complex With Nadph Chain , Alrestain Bound To C298aW219Y MUTANT HUMAN ALDOSE Reductase family 449 11 aldo-keto reductase family 1, member B10; aldose reductase-like peptide; aldose reductase-related protein; small intestine reductase family 1, member B10 (aldose reductase); aldose reductase-like peptide; aldose reductase-like lado-keto reductase family 1, member B10 (aldose reductase); aldose reductase-like lado-keto reductase family 1, member B10 (aldose reductase); aldose reductase-like lado-keto reductase family 1, member B11 (aldose reductase-like lado-keto reductase family 1, member B11 (aldose reductase related protein lado-keto reductase loopADR, aldo-keto reductase related protein lado-keto reductase family 1, member A1; aldehyde reductase, alcohol dehydrogenase family 1, member A1; aldehyde reductase, beta polypeptide 1 (3-oxo-5 297) aldo-keto reductase family 1, member B1; steroid 5-beta-reductase, beta polypeptide 1 (3-oxo-5 291) aldo-keto reductase family 1, member B1; steroid 5-beta-reductase, beta polypeptide 1 (3-oxo-5 291) aldo-keto reductase.	pdb 2ACQ	Chain, Aldose Reductase (E.C.1.1.21) Wild Type Complexed With Nadp+ And Glucose-6-Phosphate	540	1E-153
Chain , Aldose Reductase family 1, member B1 (aldose reductase) Chain , Aldose Reductase (E.C.1.1.2.1) Mutant With Cys 298 Replaced By Ser (C298s) 536 1. Complex With Nadph Chain , Aldose Reductase (E.C.1.1.1.2.1) Mutant With Cys 298 Replaced By Ser (C298s) 536 1. Chain , Altestatin Bound To C298aW219Y MUTANT HUMAN ALDOSE Reductase 533 1. Aldo-keto reductase family 1, member B10, aldose reductase-like peptide; aldose reductase-related protein; small intestine reductase family 1, member B10 (aldose reductase); aldose reductase-related protein; small intestine reductase family 1, member B10 (aldose reductase); aldose reductase-like 1; aldo-keto reductase family 1, member B10 (aldose reductase); aldose reductase family 1, member B11 (aldose reductase); aldo-keto reductase loopADR, aldo-keto reductase related protein 3.1 similar to aldose reductase loopADR; aldo-keto reductase related protein Chain , Aldeilyde Reductase Chain , Aldeilyde Reductase Chain , Aldeilyde Reductase family 1, member A1; aldeltyde reductase; alcohol dehydrogenase Chain , Aldeilyde Reductase family 1, member A1; aldethyde reductase, beta polypeptide 1 (3-oxo-5 293) beta-steroid delta 4-dehydrogenase beta 1); steroid 5-beta-reductase alde-steroid delta 4-dehydrogenase beta 1); steroid 5-beta-reductase aldehyde reductase	pdb 2ACU	Chain, Aldose Reductase (B.C.1.1.1.21) Mutant With Tyr 48 Replaced By His (Y48h) Complexed With Nado+ And Citrate	538	1E-153
Chain , Aldose Reductase (E.C.I.I.21) Mutant With Cys 298 Replaced By Ser (C298s) 536 I. Complex With Nadph Complex With Nadph Chain , Altestatin Bound To C298aW219Y MUTANT HUMAN ALDOSE Reductase 533 II. aldo-keto reductase family I, member B10; aldose reductase-like peptide; aldose reductase-related protein; small intestine reductase-like); aldose reductase-like peptide; aldose reductase-related protein; small intestine reductase family I, member B10 (aldose reductase); aldose reductase-like 1; 388 I aldo-keto reductase family I, member B11 (aldose reductase); aldose reductase-like 1; 388 I aldo-keto reductase family I, member B11 (aldose reductase-like 1; 388 I aldo-keto reductase loopADR; aldo-keto reductase related protein 310 I aldo-keto reductase loopADR; aldo-keto reductase related protein 310 Chain , Aldehyde Reductase Chain , Aldehyde Reductase family I, member A1; aldehyde reductase, alcohol dehydrogenase Chain , aldo-keto reductase family I, member A1; aldo-keto-reductase, beta polypeptide I (3-oxo-5) 293 beta-steroid delta 4-dehydrogenase beta 1); steroid 5-beta-reductase 1 aldo-keto reductase family I, member A1; aldehyde reductase, beta polypeptide I (3-oxo-5) 293 leta-steroid delta 4-dehydrogenase beta 1); steroid 5-beta-reductase 1 aldo-keto reductase 1 aldo-keto reductase family I, member A1; aldehyde reductase, aldehyde reductase family I, member A1; aldehyde reductase family I, member A1; aldehyde reductase family I, member A1; aldehyde reductase family I, member A1; aldehyde reductase family I, member A1; aldehyde reductase family I, member A1; aldehyde reductase family I, member A1; aldehyde reductase family I, member A1; aldehyde reductase family I, member A1; aldehyde reductase family I, member A1; aldehyde reductase family I, member A1; aldehyde reductase family I, member A1; aldehyde reductase family I, member A1; aldehyde reductase family I, member A1; aldehyde reductase family I, aldo-keto reductase family I, aldehyde reductase family I, aldehyde reductase family I, aldehyde II	AAH05387.1	aldo-keto reductase family 1, member B1 (aldose reductase)	537	1E-152
Chain, Alrestatin Bound To C298aW219Y MUTANT HUMAN ALDOSE Reductase Chain, Alrestatin Bound To C298aW219Y MUTANT HUMAN ALDOSE Reductase 1. aldo-keto reductase family 1, member B10; aldose reductase-like 1; aldo-kcto reductase-related protein; small intestine reductase-like); aldose reductase-like peptide; aldose reductase-related protein; small intestine reductase family 1, member B10 (aldose reductase); aldose reductase-like 1; 388 1 aldo-keto reductase family 1, member B11 (aldose reductase-like 1; 388 1 aldo-keto reductase family 1, member B11 (aldose reductase-like 1; 388 1 aldo-keto reductase loopADR; aldo-keto reductase related protein 31 aldo-keto reductase loopADR; aldo-keto reductase related protein 31 aldo-keto reductase loopADR; aldo-keto reductase related protein 31 aldo-keto reductase loopADR; aldo-keto reductase setuctase loopADR; aldo-keto reductase setuctase loopADR; aldo-keto reductase setuctase loopADR; aldo-keto reductase setuctase loopADR; aldo-keto reductase family 1, member A1; aldehyde reductase, beta polypeptide 1 (3-oxo-5 293 beta-steroid delta 4-dehydrogenase beta 1); steroid 5-beta-reductase aldehyde reductase aldehyde reductase	pdb[1ABN	Chain, Aldose Reductase (E.C.1.1.21) Mutant With Cys 298 Replaced By Ser (C298s)	536	1E-152
Chain, Altestatin Bound To C298aW219Y MUTANT HUMAN ALDOSE Reductase 533 I aldo-keto reductase family 1, member B10; aldose reductase-like 1; aldo-keto reductase family 1, member B10; aldose reductase-like); aldose reductase-like peptide; aldose reductase-related protein; small intestine reductase (AR) (Aldehyde reductase) 397 I similar to Aldose reductase family 1, member B10 (aldose reductase); aldose reductase family 1, member B10 (aldose reductase); aldose reductase family 1, member B11 (aldose reductase) 333 I aldo-keto reductase loopADR. 4.1 aldo-keto reductase loopADR; aldo-keto reductase related protein 310 Chain, Aldehyde Reductase loopADR; aldo-keto reductase reductase loopADR; aldo-keto reductase loopADR; aldo-keto reductase loopADR; aldo-keto reductase loopADR; aldo-keto reductase family 1, member A1; aldehyde reductase; alcohol dehydrogenase Chain, Aldehyde Reductase family 1, member A1; aldehyde reductase, beta polypeptide I (3-oxo-5 293 beta-steroid delta 4-dehydrogenase beta 1); steroid 5-beta-reductase aldehyde reductase aldehyde reductase		Complex With Nadph		
1.1 aldo-keto reductase family 1, member B10; aldose reductase-like 1; aldo-keto reductase family 448 1 1, member B11 (aldose reductase-like); aldose reductase-like peptide; aldose reductase-related protein; small intestine reductase. 1, member B11 (aldose reductase (AR) (Aldehyde reductase) 397 1 2.1 similar to Aldose reductase family 1, member B10 (aldose reductase); aldose reductase-like 1; 388 1 aldo-keto reductase family 1, member B11 (aldose reductase-like 1); aldo-keto reductase loopADR; aldo-keto reductase related protein 318 2.1 aldo-keto reductase loopADR; aldo-keto reductase related protein 310 Chain, Aldehyde Reductase Chain, Aldehyde Reductase family 1, member A1; aldehyde reductase, alcohol dehydrogenase 297 Chain, Aldehyde Reductase family 1, member D1; steroid-5-beta-reductase, beta polypeptide 1 (3-oxo-5) 293 beta-steroid delta 4-dehydrogenase beta 1); steroid 5-beta-reductase aldehyde reductase aldehyde reductase	pdb[1AZ1	Chain, Alrestatin Bound To C298aW219Y MUTANT HUMAN ALDOSE Reductase	533	1E-151
2 aldo-keto reductase family 1, member B10; aldose reductase-like peptide; aldo-keto reductase family 1, member B11 (aldose reductase-like); aldose reductase-like) aldose reductase-like); aldose reductase reductase (AR) (Aldehyde reductase) 2 similar to Aldose reductase family 1, member B10 (aldose reductase); aldose reductase-like 1; 388 1 aldo-keto reductase family 1, member B11 (aldose reductase-like 333 1 aldo-keto reductase loopADR; aldo-keto reductase related protein 316 reductase loopADR; aldo-keto reductase related protein 316 chain, Aldehyde Reductase loopADR; aldo-keto reductase related protein 316 chain, Aldehyde Reductase loopADR; aldo-keto reductase related protein 316 chain, Aldehyde Reductase family 1, member A1; aldehyde reductase, beta polypeptide 1 (3-oxo-5 beta-steroid delta 4-dehydrogenase beta 1); steroid 5-beta-reductase aldo-keto reductase family 1, member D1; steroid 5-beta-reductase selated selated aldehyde reductase family 1, member D1; steroid 5-beta-reductase aldehyde reductase family 1, member D1; steroid 5-beta-reductase aldehyde reductase family 291	AAC36465.1	aldo-keto reductase	449	1E-126
1, member B11 (aldose reductase-like); aldose reductase-like peptide; aldose reductase-related protein; small intestine reductase 2 similar to Aldose reductase (AR) (Aldehyde reductase) 1 similar to aldo-keto reductase family 1, member B10 (aldose reductase); aldose reductase-like 1; 2 aldo-keto reductase family 1, member B11 (aldose reductase-like 2 aldo-keto reductase loopADR 3 aldo-keto reductase loopADR; aldo-keto reductase related protein 3 aldo-keto reductase loopADR; aldo-keto reductase related protein 4 similar to aldose reductase loopADR; aldo-keto reductase related protein 5 aldo-keto reductase family 1, member A1; aldehyde reductase; alcohol dehydrogenase 6 aldo-keto reductase family 1, member A1; steroid-5-beta-reductase, beta polypeptide 1 (3-oxo-5) 8 beta-steroid delta 4-dehydrogenase beta 1); steroid 5-beta-reductase 9 aldehyde reductase	NP_064695.2	aldo-keto reductase family 1, member B10; aldose reductase-like 1; aldo-keto reductase family		1E-126
protein; small intestine reductase 2 similar to Aldose reductase (AR) (Aldehyde reductase) 2 similar to Aldose reductase family 1, member B10 (aldose reductase); aldose reductase-like 1; 388 1 aldo-keto reductase family 1, member B11 (aldose reductase-like 1 aldo-keto reductase loopADR 2 similar to aldose reductase loopADR; aldo-keto reductase related protein 3 aldo-keto reductase loopADR; aldo-keto reductase related protein 3 aldo-keto reductase loopADR; aldo-keto reductase related protein Chain , Aldehyde Reductase 3 aldo-keto reductase family 1, member A1; aldehyde reductase, beta polypeptide 1 (3-oxo-5 293 aldo-keto reductase family 1, member D1; steroid 5-beta-reductase 3 beta-steroid delta 4-dehydrogenase beta 1); steroid 5-beta-reductase 3 aldo-keto reductase 3 aldo-keto reductase family 1, member D1; steroid 5-beta-reductase, beta polypeptide 1 (3-oxo-5 291 aldo-keto reductase	 	1, member B11 (aldose reductase-like); aldose reductase-like peptide; aldose reductase-related		
2 similar to Aldose reductase (AR) (Aldehyde reductase) 1. similar to aldo-keto reductase family 1, member B10 (aldose reductase); aldose reductase-like 1; 1. aldo-keto reductase family 1, member B11 (aldose reductase-like 1. aldo-keto reductase loopADR; aldo-keto reductase related protein 2. aldo-keto reductase loopADR; aldo-keto reductase related protein 2. aldo-keto reductase loopADR; aldo-keto reductase related protein 2. aldo-keto reductase family 1, member A1; aldehyde reductase; alcohol dehydrogenase 2. aldo-keto reductase family 1, member A1; aldehyde reductase, beta polypeptide 1 (3-oxo-5 293) 2. beta-steroid delta 4-dehydrogenase beta 1); steroid 5-beta-reductase 3. aldehyde reductase		protein; small intestine reductase		
 similar to aldo-keto reductase family 1, member B10 (aldose reductase); aldose reductase-like 1; 388 aldo-keto reductase family 1, member B11 (aldose reductase-like) aldo-keto reductase family 1, member B11 (aldose reductase-like) similar to aldose reductase loopADR; aldo-keto reductase related protein aldo-keto reductase loopADR; aldo-keto reductase related protein aldo-keto reductase loopADR; aldo-keto reductase related protein aldo-keto reductase family 1, member A1; aldehyde reductase; alcohol dehydrogenase aldo-keto reductase family 1, member A1; aldehyde reductase, beta polypeptide 1 (3-oxo-5) beta-steroid delta 4-dehydrogenase beta 1); steroid 5-beta-reductase aldehyde reductase aldehyde reductase 	XP_064999.2	similar to Aldose reductase (AR) (Aldehyde reductase)	397	1E-110
aldo-keto reductase family 1, member B11 (aldose reductase-like) 1 aldo-keto reductase loopADR 4 similar to aldose reductase 1 aldo-keto reductase loopADR; aldo-keto reductase related protein 2 aldo-keto reductase loopADR; aldo-keto reductase related protein 2 aldo-keto reductase family 1, member A1; aldehyde reductase; alcohol dehydrogenase 3 aldo-keto reductase family 1, member A1; aldehyde reductase, beta polypeptide I (3-oxo-5) 3 beta-steroid delta 4-dehydrogenase beta 1); steroid 5-beta-reductase 3 aldehyde reductase	XP_069833.1	similar to aldo-keto reductase family 1, member B10 (aldose reductase); aldose reductase-like 1;		1E-108
 1 aldo-keto reductase loopADR. 3 similar to aldose reductase 3 similar to aldose reductase 3 aldo-keto reductase loopADR; aldo-keto reductase related protein 3 aldo-keto reductase loopADR; aldo-keto reductase related protein 3 aldo-keto reductase family 1, member A1; aldehyde reductase; alcohol dehydrogenase 3 aldo-keto reductase family 1, member A1; steroid-5-beta-reductase, beta polypeptide 1 (3-oxo-5 aldehyde reductase alidehyde reductase 3 beta-steroid delta 4-dehydrogenase beta 1); steroid 5-beta-reductase 3 aldehyde reductase 3 aldehyde reductase 3 aldehyde reductase 		aldo-keto reductase family 1, member B11 (aldose reductase-like		
 similar to aldose reductase 1 aldo-keto reductase loopADR; aldo-keto reductase related protein 1 aldo-keto reductase loopADR; aldo-keto reductase related protein 297 1 aldo-keto reductase family 1, member A1; aldehyde reductase; alcohol dehydrogenase 1 aldo-keto reductase family 1, member D1; steroid-5-beta-reductase, beta polypeptide 1 (3-oxo-5 beta-steroid delta 4-dehydrogenase beta 1); steroid 5-beta-reductase 291 292 293 294 294 296 296 296 296 291 	AAK58523.1	aldo-keto reductase loopADR	333	4E-91
 aldo-keto reductase loopADR; aldo-keto reductase related protein aldo-keto reductase loopADR; aldo-keto reductase related protein Chain, Aldehyde Reductase aldo-keto reductase family 1, member A1; aldehyde reductase; alcohol dehydrogenase aldo-keto reductase family 1, member D1; steroid-5-beta-reductase, beta polypeptide 1 (3-oxo-5 beta-steroid delta 4-dehydrogenase beta 1); steroid 5-beta-reductase aldehyde reductase 	XP_089195.4	similar to aldose reductase	332	1E-90
1 aldo-keto reductase loopADR; aldo-keto reductase related protein Chain , Aldehyde Reductase Chain , Aldehyde Reductase 1 aldo-keto reductase family 1, member A1; aldehyde reductase; alcohol dehydrogenase 1 aldo-keto reductase family 1, member D1; steroid-5-beta-reductase, beta polypeptide 1 (3-oxo-5 293 beta-steroid delta 4-dehydrogenase beta 1); steroid 5-beta-reductase 297 297 298 299	NP_113624.1	aldo-keto reductase loopADR; aldo-keto reductase related protein	318	1E-86
Chain, Aldehyde Reductase Chain, Aldehyde Reductase 1 aldo-keto reductase family 1, member A1; aldehyde reductase; alcohol dehydrogenase 297 1 aldo-keto reductase family 1, member D1; steroid-5-beta-reductase, beta polypeptide 1 (3-oxo-5 293 beta-steroid delta 4-dehydrogenase beta 1); steroid 5-beta-reductase 297 298 299	XP_166652.1		310	4E-84
1 aldo-keto reductase family 1, member A1; aldehyde reductase; alcohol dehydrogenase 297 aldo-keto reductase family 1, member D1; steroid-5-beta-reductase, beta polypeptide 1 (3-oxo-5 293 beta-steroid delta 4-dehydrogenase beta 1); steroid 5-beta-reductase 291	pdb 2ALR		297	3E-80
293	NP_006057.1		297	3E-80
beta-steroid delta 4-dehydrogenase beta 1); steroid 5-beta-reductase	NP_005980.1	aldo-keto reductase family 1, member D1; steroid-5-beta-reductase, beta polypeptide 1 (3-oxo-5		5E-79
aldehyde reductase , 291		beta-steroid delta 4-dehydrogenase beta 1); steroid 5-beta-reductase		
	1403439A	aldehyde reductase	291	2E-78

•			252		
		BAA99542.1	A99542.1 3alpha-hydroxysteroid dehydrogenase variant	287	3E-77
		AAH20744.1	aldo-keto reductase family I, member C4 (chlordecone reductase; 3-alpha hydroxysteroid dehydrogenase, type I; dihydrodiol dehydrogenase 4)	286	4E-77
		NP_001809.1	NP_001809.1 aldo-keto reductase family 1, member C4 (chlordecone reductase; 3-alpha hydroxysteroid dehydrogenase, type I; dihydrodiol dehydrogenase 4); chlordecone reductase; type I 3-alpha-HSD. Chlordecone reductase	286	4E-77
		BAA05122.1	3 alpha-hydroxysteroid/dihydrodiol dehydrogenase DD4	286	4E-77
		NP_001344.2	NP_001344.2 aldo-keto reductase family 1, member Cl; dihydrodiol dehydrogenase 1; dihydrodiol dehydrogenase; trans-1,2-dihydrobenzene-1,2-diol dehydrogenase; chlordecone reductase homolog; 20 alpha-hydroxysteroid dehydrogenase; aldo-keto reductase C; hepatic dihydrodiol dehydrogenase	283	5E-76
		AAA35658.1	AAA35658.1 chlordecone reductase	280	4E-75
NM_013484 Mm.2081	F;2.02 (5to19)	NP_000054.2	_000054,2 complement component 2 precursor; C3/C5 convertase	1136	0
NP 038512.1		С2НО	complement C2 precursor [validated]	1134	0
		BAB63292.1	C2	873	0
		NP_001701.1	complement factor B preproprotein; B-factor, properdin; C3 proactivator; C3 proaccelerator; glycine-rich beta-glycoprotein; C3/C5 convertase	495	1E-139
		P00751	Complement factor B precursor (C3/C5 convertase) (Properdin factor B) (Glycine-rich beta glycoprotein) (GBG) (PBF2)	495	1E-139
		AAH04143.1	B-factor, properdin	495	1E-139
		AAA36225.2	MHC serum complement factor B	427	1E-119
	-	AAH29781.1	Similar to complement component 2	412	1E-114

Ľ	٦
C	Į

1E-95	7 1E-104) 2E-97	5 2E-96	1 3E-93		3E-84		7 2E-86		0 2E-51	0	0	0	0	0	0	6 1E-146	
349	377		269	265	341	_	310	- <u></u>	317		200	1511	1509	1507	1499	1498	1327	516	
factor B	612382.1 hypothetical protein BC013995		unnamed protein product	similar to hypothetical protein BC013995	unnamed protein product		036323.1 putative tumor suppressor FUS2		glutathione peroxidase 6		unnamed protein product	mannosidase, alpha, class 2B, member 1	mannosidase, alpha, class 2B, member 1; mannosidase, alpha B, lysosomal	lysosomal alpha-mannosidase	alpha-mannosidase	.C50812.1 lysosomal acid alpha-mannosidase	alpha-mannosidase (EC 3.2.1.24) precursor	truncated lysosomal acid alpha-mannosidase	•
AAK30167.1 factor B	NP_612382.1		BAB71502.1	XP_065813.2	BAC04265.1		NP_036323.1		NP_056511		BAB85019	AAH00736.1	NP_000519.1	AAC34130.1	AAB03816.1	AAC50812.1	JC2200	AAC50811.1	
	F:2.02 (5to19)						F:2.02 (5to19)		F:2.01 (YtoO)			F:2.01 (7to19)							
							I												
	NM_016969 Mm.29874	NP 058665.1	,				NM_019750 Mm.29271	NP 062724.1	NM_024198 Mm.20164	NP 077160.1		NM_010764 Mm.4219 NP_033268 1							

Ŋ

C)

			254		
			2)		
		Q9Y2E5	Epididymis-specific alpha-mannosidase precursor (Mannosidase alpha class 2B member 2)	294	6E-79
12 Mm.12604	F:2.01 (7to19)	NP_036370.2	type information regulation 2, S. cerevisiae, homolog) 1; sirtuin	1032	0
NP 062786.1			type 1; sir2-like 1; SIR2alpha		
		CAC04174.1	bA57G10.4 (SIRT1, Sir2-like proteins (siruitins) type 1)	934	-
		AAH12499.1	Unknown (protein for MGC:21066)	928	0
B C 0 0 6 6 2 1 Mm.28488 AAH06621.1	F:2.01 (5to11)	NP_055764.1	_055764.1 KIAA0907 protein	752	0
		AAH27182.1	Sinular to KIAA0907 protein	319	2E-85
0.2	F:2.01 (5to19)	AAH09503.1	G1 to S phase transition 1	872	0
BAA32526.1					
		NP_002085.1	NP_002085.1 G1 to S phase transition 1	898	c
		CAB91089.1	CAB91089.1 polypeptide chain release factor 3b	861	0
		NP_060564.1	060564.1 peptide chain release factor 3	860	0
		AAH36077.1	G1 to S phase transition 2	855	0
		BAB14435.1	unnamed protein product	962	0
		XP_017821.2	017821.2 similar to peptide chain release factor 3	354	2E-96
		BAA82990.1	KIAA1038 protein	316	7E-85

2.5

7E-85	3E-55		1E-155			4E-86	1E-85	 3E-84	3E-67	2E-65	9E-65	2E-64	4E-58	SE-55	5E-55	1E-54			4F-54
316 7E	1		551 1E-			320 4E	318 1H	 313 3I	257 3I	251 2I	249 91	248 21	227 41	216 51	216 5	215 1]			213 4
	NP 076869 1 hymothetical protein IMAGE3455200		NP_000157.1 gap junction protein, beta 1, 32kDa (connexin 32, Charcot-Marie-Tooth neuropathy, X-linked); 55	Gap junction protein, beta-1, 32kD (connexin 32); gap junction protein, beta 1, 32kD (connexin	32, Charcot-Marie-Tooth neuropathy, X-linked)		gap junction protein, beta 2, 26kDa (connexin 26); gap junction protein, beta 2, 26kD (connexin) 31	junction protein, beta 6 (connexin 30)	similar to Gap junction beta-4 protein (Connexin 30.3) (Cx30.3)	25259.1 gap junction protein, beta 5 (connexin 31.1)		junction protein, beta 3, 31kDa (connexin 31); gap junction protein, beta 3, 31kD (connexin	nexin25	Gap junction alpha-8 protein (Connexin 50) (Cx50) (Lens fiber protein MP70)	Da liver protein)	005258.1 gap junction protein, alpha 8, 50kDa (connexin 50); gap junction membrane channel protein 21	alpha-8; connexin 50; Gap junction membrane channel protein alpha-8 (connexin 50); gap	junction protein, alpha 8, 50kD (connexin 50)	068773 2 Jan innetion protein alpha 3 46kDs (connevin 46) can innetion protein alpha 3 46kD 21
NP_006611.1 HBS1-like	NP 076869 1 hv			Ga	32,	AAF91440.1 gap	NP 003995.1 gai	NP_006774.1 gal	XP_060532.1 sin	NP_005259.1 ga	AAC95472.1 connexin 31.1	NP_076872.1 ga	CAC93845.1 con	P48165 Ga	XP 088689.1 sir	NP_005258.1 ga	alg	mi.	1 CCC 070 atk
	E-2 ()1 (5to 19)	(2002) 10:2:1	F:2.01 (5to19)																
	T		1																
	N V 0 0 3 2 3 7 Mm 20703	BAB22661.1	NM_008124 Mm.21198	NP_032150.2															

10

NM_008723 Mm.1406 F:2.01 (5to19)	76H8	IGan iunction alpha-3 protein (Connexin 46) (Cx46)	713	4E-54
23 Mm.1406		The state of the s	417	ナレーゴナ
23 Mm.1406	NP_000156.1	connexin 43; gap junction protein, alpha 1, 43kD (connexin 43); gap junction protein, alpha 1,	203	6E-51
2 3 Mm.1406		43kD		
23 Mm.1406				
NP 032749 1	g N	008924.1 nucleophosmir/nucleoplasmin 3; nucleoplasmin-3; nucleophosmin/nucleoplasmin family, member	234	2E-61
11717		3		
	AAD51496.1	AAD51496, 1 nucleophosmin/nucleoplasmin3	224	1E-58
NM_022325 Mm.156919 F:2 (7to19) NP 071720.1	XP	030699.1 similar to Cathepsin Z precursor (Cathepsin X) (Cathepsin P)	539	1B-153
	AAC39839.1	cathepsin Z precursor; CTSZ	539	IE-153
	AAC63141.1	preprocathepsin P	539	1B-153
	pdb 1DEU	Chain A, Crystal Structure Of Human Procathepsin X: A Cysteine Protease With The Proregion	536	1E-152
		Covalently Linked To The Active Site Cysteine		
	AAC61477.1	cathepsin X precursor	534	1E-151
	pdb 1EF7	Chain A, Crystal Structure Of Human Cathepsin X	474	1E-133
M 6 2 3 6 1 Mm.1779 F.2 (5to19)	ď.	_002970.2 sterol carrier protein 2	390	1E-107
AAA40099.1				
	AAA03558.1	sterol carrier protein-2	390	1E-107
	AAB41286.1	sterol carrier protein-X/sterol carrier protein-2	387	1E-107
	B40407	sterol carrier protein 2-related form, 58.85K	378	1E-104
	AAA03559.1	AAA03559.1 sterol carrier protein-2	245	4E-64
	AAB24921.1	sterol carrier protein 2; SCP2	236	2E-61
	DAP 1QND	Chain A, Sterol Carrier Protein-2, Nrnt, 20 Structures	208	7E-53

Ŋ

2

7)

_	ĺ
Ŋ	Į
C	Ì
	۱

				100		
NM_008331 Mm.6718 F:2 (5to19)	Mm.6718	F:2 (5to19)	AAH07091.	Unknown (protein for MGC:14710)	456	456 1E-127
NP 032357.1						
			NP_001539.1	001539.1 interferon-induced protein with tetratricopeptide repeats 1; Interferon, alpha-inducible protein	455	1E-126
				(MW 56kD); interferon-induced protein 56		
			NP_036552.1	036552.1 retinoic acid- and interferon-inducible protein (58kD)	395	1E-108
			XP_084477.1	084477.1 similar to Interferon-induced protein with tetratricopeptide repeats 2 (IFIT-2) (Interferon-induced	295	1E-78
				54 kDa protein) (IFI-54K) (ISG-54 K)		
			AAH32839.1	Similar to interferon-induced protein with tetratricopeptide repeats 2	274	3E-72
			NP_001540.1	NP_001540.1 interferon-induced protein with tetratricopeptide repeats 4	272	2E-71
			XP_048183.1	similar to Interferon-induced protein with tetratricopeptide repeats 4 (IFIT-4) (Interferon-induced	271	3E-71
				60 kDa protein) (IFI-60K) (ISG-60) (CIG49) (Retinoic acid-induced gene G protein) (RIG-G)		
	_		AAH04977.1	AAH04977.1 Interferon-induced protein with tetratricopeptide repeats 4	268	2E-70

ហ

Table 1B: Unfavorable Genes/Proteins	Genes/Protein	SI			
Mouse Gene Unigene	Behavior	Human	n Description	Bits	B
Protein		Proteins	4		
NM_007702 Mm.449	U:52.77 (YtoO)	AAC34987.1	AAC34987.1 cell death activator CIDE-A	340	3E-92
INF 031/20.1		AAH31896.1	AAH31896.1 Similar to cell death-inducing DFFA-like effector a	319	5E-86
NM_007822 Mm.7459	U:18.8 (5to7)	NP_000769.1	NP_000769.1 cytochrome P450, subfamily IVA, polypeptide 11; fatty acid omega-hydroxylase; P450HL-	780	0
NP 031848.1	-		omega; alkane-1 monooxygenase; lauric acid omega-hydroxylase		
		002928	Cytochrome P450 4A11 precursor (CYPIVA11) (Fatty acid omega-hydroxylase) (P-450 HK	LLL	0
`	,		omega) (Lauric acid omega-hydroxylase) (CYP4AII) (P450-HL-omega)		
-		165981	fatty acid omega-hydroxylase (EC 1.14.15) cytochrome P450 4A11 AAB80170, Sequence 6	765	0
			from patent US 5667992		
	 - -	BAA02864.1	BAA02864.1 fatty acid omega-hydroxylase	19/	0
		1 0007001	Part 13 1 America	ì	1
	***************************************	AAr /6/22.1	iatty acid omega-nydroxylase CYP4A11	746	0
		CAB72105.1	CAB72105.1 dJ18DJ4.4 (cytochrome P450, subfamily IVA, polypeptide 11)	736	0
		O4HUB1	O4HUB1 cytochrone P450 4B1	499	1E-139
			NP_505847 Cytochrome P450 family member (57.2 kD) [Caenorhabditis elegans]		
		AAL57720.1	cytochrome P450	499	IE-139
		AAM09532.1	AAM09532.1 cytochrome P450	499	1E-139
		NP_000770.1	NP_000770.1 cytochrome P450, subfamily IVB, polypeptide 1; cytochrome P450, subfamily IVB, member 1;	497	1E-139
			microsomal monooxygenase		
		AAL57721.1	cytochrome P450	497	1E-139
		AAH17758.1	AAH17758.1 Unknown (protein for MGC:22150)	495	1E-138
		AAH28102.1	AAH28102.1 Unknown (protein for MGC:40051)	489	1E-137

ப

Ŋ

 $\ddot{\sim}$

			100		
		BAC03751.1	C03751.1 unnamed protein product	448	1E-124
		BAC04868.1	BAC04868.1 unnamed protein product	405	1E-110
		BAA75823.1	BAA75823.1 Leukotriene B4 omega-hydroxylase	398	1E-109
		NP_001073.3	001073.3 cytochrome P450, subfamily IVF, polypeptide 2; leukotriene B4 omega-hydroxylase; leukotriene-	398	1E-109
			B4 20-monooxygenase		
		NP_000887.1	000887.1 cytochrome P450, subfamily IVF, polypeptide 3; leukotriene B4 omega hydroxylase; leukotriene-	394	1E-108
			B4 20-monooxygenase; cytochrome P450-LTB-omega		
		AAC50052.2	AAC50052.2 cytochrome P450 4F2	393	1E-108
		AAC08589.1	AAC08589.1 cytochrome P-450	390	1E-107
		Q9HBI6	Cytochrome P450 4F11 (CYPIVF11)	387	1E-106
		NP_067010.1	cytochrome P450, subfamily IVF, polypeptide 11	387	1E-106
		Q9HCS2	Q9HCS2 Cytochrome P450 4F12 (CYPIVF12)	384	1E-105
		NP_076433.1	cytochrome P450 isoform 4F12	384	1E-105
		AAH35350.1	AAH35350.1 similar to cytochrome P450	384	1E-105
		AAC11543.1	F22329_1	381	1E-104
		NP_009184.1	NP_009184.1 cytochrome P450, subfamily IVF, polypeptide 8; microsomal monooxygenase; flavoprotein-linked	380	1E-104
			топоохудепаse		
		CAD38795.1	AD38795.1 hypothetical protein	347	6E-94
	-	XP_065069.2	XP_065069.2 record was removed	337	6E-91
		XP_029070.2	XP_029070.2 record was removed	323	9E-87
		AAH22851.1	Similar to cytochrome P450, subfamily IVA, polypeptide 11	287	6E-76
		XP_065068.1	XP_065068.1 record was removed	278	2E-73
		BAC05026.1	unnamed protein product	278	2E-73
		BAA02145.1	BAA02145.1 cytochrome P-450LTBV	270	6E-71
		CAA50586.1	CAA50586.1 cytochrome P450	263	1E-68
		AAL57719.1	AAL57719.1 truncated cytochrome P450	237	5E-61
NM_008745 Mm.3993	U;14,81 (YtoO) AAM77876		protein tyrosine kinase non catalytic form	898	0
NP 032771.1					
·		AAL67965	neurotrophin receptor tyrosine kinase type 2	846	0

10

Ci

			260		
		AAL67967	neurotrophin receptor tyrosine kinase type 2 truncated isoform	846	0
		AAL67966	neurotrophin receptor tyrosine kinase type 2 truncated isoform	845	0
		A56853	brain-derived neurotrophic factor receptor precursor	845	0
•		CAA12029	TRKC	273	1E-71
		2103287B	trkC gene	273	1E-71
		AAH13693	Unknown (protein for MGC:17113)	273	1E-71
		12	trkC	273	1E-71
		A55178	neurotrophin receptor trkC precursor	273	1E-71
		BAA34355	TRKA	237	7E-61
		P04629	High affinity nerve growth factor receptor precursor (TRK1 transforming tyrosine kinase protein)	236	1E-60
			(p140-TrkA) (Trk-A).		
		AAA36770	trk tyrosine-specific protein kinase	229	2E-58
		1WWBX	Chain X, Ligand Binding Dornain Of Human Trkb Receptor	223	1E-56
		1HCFX	Chain X, Crystal Structure Of Trkb-D5 Bound To Neurotrophin-45.	216	2E-54
	•				
NM_026574 Mm.89148	U:12.76 (5to11)	BAA86573.1	BAA86573.1 KIAA1259 protein	775	0
NP 080850.1					
		BAA92122.1	unnamed protein product	364	6E-99
		T46350	hypothetical protein DKFZp434B0616.1 - human	268	4E-70
NM_021456 Mm.22720	U:10.66 (YtoM) AAH12418	AAH12418	Unknown (protein for MGC:9220)	298	0
NP 067431.1				,	
		NP_001257	carboxylesterase 1 (monocyte/macrophage serine esterase 1); liver carboxylesterase;	865	0
			carboxylesterase 2 (liver)		
		AAA35711	carboxylesterase.	863	0
		BAB85656	brain carboxylesterase hBr2	863	0
		BAA04650	carboxylesterase	863	0
		BAA84996	brain carboxylesterase hBr3	863	0
		A41010	carboxylesterase (EC 3.1.1.1) precursor, monocyte/macrophage [validated] - human	862	0
		AAC60631	acyl coenzyme A:cholesterol acyltransferase	857	0

		A A D 52175	Mischer Charles and Charles an	857	0
		011000	სწმაჭი		ľ
		AAA16036	carboxylesterase	854	Э
		AAA35650	carboxylesterase	772	0
		CAA37147	serine esterase N-terminal truncated (503 AA)	692	0
		AAA83932	carboxylesterase	655	0
		BAA84995	brain carboxylesterase hBr1	637	0
		AAH32095	Similar to carboxylesterase 2 (intestine, liver)	468	1E-130
		AAB03611	carboxylesterase hCE-2	468	1E-130
		CAA70831	carboxylesterase	468	1E-130
		CAD28531	hypothetical protein	457	1E-127
		AAF14185	carboxylesterase-related protein	392	1E-108
		BAB71094	unnamed protein product	369	1E-101
		BAC03565	unnamed protein product	360	4E-98
		NP_776176	hypothetical protein FLJ37464	283	1E-74
		XP_208869	similar to carboxylesterase 1 (monocyte/macrophage serine esterase 1); liver carboxylesterase;	280	5E-74
	•		carboxylesterase 2 (liver)		
NM_013641 Mm.4501	U:8.87 (YtoM)	A49690	prostaglandin E receptor, subtype EP1	408	1E-112
NP 038669.1					
-		BAC05723	seven transmembrane helix receptor	322	2E-86
AK004768 Mm.31024	U:7.6 (YtoO)	BAA31679.2	BAA31679.2 KIAA0704 protein	1435	0
		NP 663160	oxysterol-binding protein-like protein 3 isoform b; oxysterol-binding protein-related	1432	0
		l 	rotein 3		
		AAM27386	oxysterol binding protein-related protein 3 isoform 1a	1416	0
		AAM27389	oxysterol binding protein-related protein 3 isoform 1 d	1353	0
		NP_663161	oxysterol-binding protein-like protein 3 isoform c; oxysterol-binding protein-related protein 3;	1337	0
			oxysterol-binding protein 3; OSBP-related protein 3		

10 20 20 25

		202		
		OSBP-related protein 6; ORP6	914	0
-	NP_665682	oxysterol-binding protein-like protein 6 isoform b;oxysterol-binding protein-related protein 6;	\$68	0
		OSBP-related protein 6		
	BAC04248	unnamed protein product	894	0
	NP_663164	oxysterol-binding protein-like protein 3 isoform f;oxysterol-binding protein-related protein 3;	828	0
		oxysterol-binding protein 3; OSBP-related protein 3		
	NP_663163	oxysterol-binding protein-like protein 3 isoform e;oxysterol-binding protein-related protein 3;	842	0
-		oxysterol-binding protein 3; OSBP-related protein 3		
	BAB55223	unnamed protein product	733	0
	AAG53410	OSBP-related protein 7; ORP7	733	0
	AAL40659	oxysterol-binding protein-like protein OSBPL7	721	0
	AAB83939	OXYSTEROL-BINDING PROTEIN; 45% similarity to P22059	423	1E-117
,	AAG53408	OSBP-related protein 3; ORP3	396	1E-109
	AAM74166	oxysterol binding protein-related protein 3 isoform 2d	386	1E-105
	AAM74165	oxysterol binding protein-related protein 3 isoform 2c	370	1E-101
	BAA91043	unnamed protein product	306	2E-81
	AAC26985	match to EST AA376471 (NID:g2028790)	299	2E-79
·	NP_542164	oxysterol-binding protein-like 1A isoform B; oxysterol-binding protein-related protein 1;	569	2E-70
		oxysterol-binding protein-like 1B; OSBP-related protein 1		
	AAL40663	oxysterol-binding protein-like protein OSBPL1B	569	2E-70
	AAL40662	oxysterol-binding protein-like protein OSBPL1A	569	2E-70
	AAK15154	oxysterol-binding protein-related protein	569	2E-70
	CAC22307	bA157P1.3.1 (KIAA0772, isoform 1)	263	9E-69
	AAL40660	oxysterol-binding protein-like protein OSBPL2	263	9E-69
	AAH11581	Similar to oxysterol binding protein	261	2E-70
	BAB33334	KIAA1664 protein	246	2E-63
	Q969R2.	Oxysterol-binding protein 2 (Oxysterol binding protein-related protein 4) (OSBP-related protein	246	2E-63
		4) (ORP-4).		
	AAG53406	OSBP-related protein 4	246	2E-63

R.

10

H

WO 2005/000335 PCT/US2004/017322

263

			263		
			unnamed protein product	246	2E-63
		AAC12953	OXYSTEROL-BINDING PROTEIN-like; similar to P22059 (PID:g129308)	242	2E-70
		AAC26986	similar to oxysterol-binding proteins; 76% Similarity to P22059	241	2E-70
		NP_579802	oxysterol-binding protein-like 1A isoform C, oxysterol-binding protein-related protein 1;	222	2E-70
			oxysterol-binding protein-like 1B; OSBP-related protein 1		
		BAA91496	unnamed protein product	216	2E-70
AK011986 Mm.40657	57 U:6.54 (7to19)	NP_653290	hỳpothetical protein FLJ32191	325	1E-87
BAB27959.1					
		NP_659448	zinc finger protein 25	323	7E-87
·		BAB71272	unnamed protein product	313	7E-84
NM_020568 Mm.12966	56 U:6.5 (YtoO)	BAB67774.1	BAB67774.1 KIAA1881 protein	1524	0
NP 065593.1					
		XP_170901.1	XP_170901.1 similar to KIAA1881 protein	248	6E-64
NM_013459 Mm.4407 NP 038487.1	7 U:6.09 (5to11)	P00746	Complement factor D precursor (C3 convertase activator) (Properdin factor D) (Adipsin)	370	1E-102
		CAC48304.1	AC48304.1 adipsin/complement factor D precursor	358	4E-99
		67580	complement factor D (EC 3.4.21.46) precursor [validated] - luman (fragment)	352	5E-97
		6730437	Chain A, Proenzyme Of Human Complement Factor D, Recombinant Profactor D	340	1E-93
		1633237	Chain, Mutant Of Factor D With Enhanced Catalytic Activity	330	1E-90
		5542120	5542120 Chain, Human Complement Factor D In Complex With Isatoic Anhydride Inhibitor	329	3E-90
		XP_084037.1	similar to Complement factor D precursor (C3 convertase activator) (Properdin factor D)	328	8E-90
			(Adipsin)		
		NP_001919.1	NP_001919.1 adipsin/complement factor D precursor	324	1E-88
NM_008182 Mm.197422 U:5.76(5to19)	422 U:5.76 (5to19)	NP_665683.1	NP_665683.1 glutathione S-transferase A1; GST, class alpha, 1; glutathione S-alkyltransferase A1; glutathione	328	8E-90
NP 032208.1			S-aryltransferase A1; S-(hydroxyalkyl)glutathione lyase A1; glutathione S-aralkyltransferase A1;		

110 20 20

				101		
				GST-epsilon; glutathione S-transferase 2		
			152381	record was removed 152381:GI number 2473582 references a Nucleotide record; you are	327	1E-89
		,		currently using the Protein database		
	!		DAA00071.1	A00071.1 TPA: glutathione transferase A5	327	1E-89
			442977	Chain A, Glutathione S-Transferase A1-1 (E.C.2.5.1.18)	326	3E-89
			1127144	Chain A, Glutathione Transferase A1-1 Complexed With An Ethacrynic Acid Glutathione	325	6E-89
			:1	Conjugate (Mutant R15k)		
			XP_167100.2	XP_167100.2 record was removed	325	6E-89
			Q1 <i>677</i> 2	Glutathione S-transferase A3-3 (GST class-alpha)	324	1E-88
			NP_000838.2	000838.2 record was removed	322	3E-88
			A49365	GI number 2302844 references a Nucleotide record; you are currently using the Protein database.	322	4E-88
,			AAA74634.1	AAA74634.1 glutathione S-transferase A3	322	4E-88
			S20331	record was removed	318	5E-87
			S27110	record was removed	317	1E-86
			S24330	glutathione transferase (EC 2.5.1.18) alpha-2 (clone GTH2)	316	3E-86
			NP_000837.2	NP_000837.2 glutathione S-transferase A2; glutathione S-transferase 2; GST, class alpha, 2; liver GST2;	315	4E-86
				glutathione S-alkyltransferase A2; glutathione S-aryltransferase A2; S-(hydroxyalkyl)glutathione		
		-,		lyase A2; glutathione S-aralkyltransferase A2; GST-gamma; HA subunit 2		
			CAB92770.1	B92770.1 dJ152L7.3 (glutathione S-transferase A2)	315	4E-86
			877958	glutathione transferase (EC 2.5.1.18) alpha-2 (clone GTH2 (+))	309	3E-84
· ·			A56801	record was removed A56801 GI number 3712816 references a Nucleotide record, you are	309	4E-84
				currently using the Protein database.		
NM_009381 Mm.28585	28585	U:5.69 (YtoO)	CAA69685	Spot14 protein	221	3E-57
NP 033407.1						
-						
AK016553 Mm.7	Mm.75856	U:5.55 (YtoO)	NP_008962	heat shock transcription factor 2 binding protein; heat shock factor 2 binding protein	549	1E-155
BAB30300.1			Ī			
			BAA95539	heat shock transcription factor 2 binding protein	378	1E-103

30

മ

4

١	Ω
C	V

				765	•	
U89406	_	U:5.43 (YtoO)	NP_004095.	fatty acid synthase	270	9E-72
AAC36513.1						
			AAH07267	Unknown (protein for IMAGE:3138929)	270	9E-72
			G01880	fatty-acid synthase (EC 2.3.1.85) (version 2)	797	4E-71
			A57788	enoyl-[acyl-carrier-protein] reductase (NADPH2, B-specific) (EC 1.3.1.10) (version 1)	254	3E-67
NM_025541 Mm.28606		U:5.13 (YtoM)	CAB43363	hypothetical protein	414	1E-114
NP 079817.1						-
			AAF29110	HSPC146	412	1E-114
			CAC05410	dJ329L24.2 (hypothetical 23.0 KD protein.)	339	6E-92
			AAK82973	anti-silencing function 1B	290	3E-77
			BAA91602	unnamed protein product	289	8E-77
A F 2 8 1 0 4 5 Mm.87471	ŀ	U:4.86 (5to11)	NP_066956.1	066956.1 ribonuclease L (2',5'-oligoisoadenylate synthetase-dependent); ribonuclease 4	904	0
AAG33708.1				-		
			A45771	A45771	006	0
A K 0 0 6 0 9 6 Mm.38305		U:4.75 (YtoO)	AAH11587	Similar to RIKEN cDNA 1700018018 gene	179	0
BAB24407.1						
			BAC04100	unnamed protein product	770	0
			AAH06353	Similar to RIKEN cDNA 1700018O18 gene	955	1E-155
NM_008495 Mm.43831	3831	U:4.6 (7to11)	NP_002296.1	002296.1 beta-galactosidase binding lectin precursor; Lectin, galactose-binding, soluble, 1;	259	2E-69
NP 032521.1						
-			1713410A	beta galactoside soluble lectin	257	6E-69
NM_025429 Mm.46316		U:4.44 (5to19)	NP_109591.1	NP_109591.1 serine (or cysteine) proteinase inhibitor, clade B (ovalbumin), member 1; protease inhibitor 2	<u>L</u>	498 1E-140
NP_079705.1		1		(anti-elastase), monocyte/neutrophil; protease inhibitor 2 (anti-elastase), monocyte/neutrophil		
	7			Juetived		

				266	L	
•.			NP_004146.1	NP_004146.1 serine (or cysteine) proteinase inhibitor, clade B (ovalbumin), member 9; protease inhibitor 9	276	6E-74
				(ovalbumin type)		
	1		NP_005015.1	serine (or cysteine) proteinase inhibitor, clade B (ovalbumin), member 10; protease inhibitor 10	275	1E-73
				(ovalbumin type, bomapin)		
			NP_002631.1	NP_002631.1 serine (or cysteine) proteinase inhibitor, clade B (ovalbumin), member 8; protease inhibitor 8	275	2E-73
				(ovalbumin type)		
			NP_004559.2	004559.2 serine (or cysteine) proteinase inhibitor, clade B (ovalbumin), member 6; protease inhibitor 6	272	1E-72
				(placental thrombin inhibitor)		
			A48681	placental thrombin inhibitor - human	569	9E-72
			138202	leupin precursor - human	267	3E-71
			XP_036951.4	036951.4 similar to Squamous cell carcinoma antigen 2 (SCCA-2) (Leupin	266	8E-71
			15988197	Chain A, Human Plasminogen Activator Inhibitor-2. [loop (66-98) Deletionmulant] Complexed With Peptide Mimicking The Reactive Center Loop	265	2E-70
			2118383	squamous cell carcinoma antigen 1 - human	264	3E-70
A F 3 3 2 0 5 2 Mm.25316	m.25316	U:4.08 (YtoO)	AAH06195	ATP citrate lyase	2101	0
AAK56081.1						
			AAB60340	ATP:citrate lyase.	2100	0
			CAA45614	ATP-citrate (pro-S-)-lyase	2046	0
			BAC04484	unnamed protein product	1340	0
A K 0 1 8 2 2 6 Mm.92685	m.92685	U:4.01 (5to19)	NP_109591.1	NP_109591.1 serine (or cysteine) proteinase inhibitor, clade B (ovalbumin), member 1; protease inhibitor 2	345	1E-138
XP_181363.1				(anti-elastase), monocyte/neutrophil; protease inhibitor 2 (anti-elastase), monocyte/neutrophil		
				derived		
			NP_004146.1	004146.1 serine (or cysteine) proteinase inhibitor, clade B (ovalbumin), member 9; protease inhibitor 9	200	5E-79
				(ovalbumin type)		
3			NP_002631.1	002631.1 serine (or cysteine) proteinase inhibitor, clade B (ovalburnin), member 8, protease inhibitor 8	207	2E-76
				(ovalbumin type)		

Ŋ

10

Á,,

				267		
			NP_005015.1	P_005015.1 serine (or cysteine) proteinase inhibitor, clade B (ovalburnin), member 10; protease inhibitor 10	179	4E-75
				(ovalbumin type, bomapin)		
			NP_004559.2	NP_004559.2 serine (or cysteine) proteinase inhibitor, clade B (ovalbumin), member 6; protease inhibitor 6	192	4E-75
				(placental thrombin inhibitor)		
			15988197	Chain A, Human Plasninogen Activator Inhibitor-2. [loop (66-98) Deletionmutant] Complexed	199	5E-75
				With Peptide Mimicking The Reactive Center Loop		
NM_010831 Mm.23789	Mm.23789	U:3.91 (YtoO)	XP_097818	similar to Probable serine/threonine protein kinase SNF1LK	1172	0
NP 034961.1						
			NP_775490	SNF1-like kinase	1171	0
		·	BAA95536	gene similar to rat protein kinase (KID2)	1163	0
			BAA34501	KIAA0781 protein	572	1E-161
			XP_041314	similar to Probable serine/threonine protein kinase SNF1LK	572	1E-161
			BAB91442	KIAA0781 protein	512	1E-143
			BAA76843	KIAA0999 protein	412	1E-113
			AAH08771	Similar to ELKL motif kinase	360	1E-97
				Ser/Thr protein kinase PAR-1Balpha	360	1E-97
			NP_004945	MAP/microtubule affinity-regulating kinase 2 isoform b; ELKL motif kinase 1; ELKL motif kinase kinase	359	2E-97
			G01025	serine/threonine protein kinase	359	2E-97
			NP_059672	MAP/microtubule affinity-regulating kinase 2 isoform a; ELKL motif kinase 1; ELKL motif kinase	359	2E-97
			NP_061120	MAP/microtubule affinity-regulating kinase 1	357	1E-96
			AAC15093	Cdc25C associated protein kinase C-TAK1	352	2E-95
			AAH24773	Unknown (protein for MGC:29880)	352	2E-95

			268		
		VAL23683	MARK4 serinc/thrconinc protein kinasc	352	2E-95
		AAK82367	Ser/Thr protein kinase PAR-1A	352	2E-95
		BAB47489	KIAA1860 protein	352	2E-95
			probable serine/threonine-specific protein kinase (EC 2.7.1)	351	5E-95
			MAP/microtubule affinity-regulating kinase 3 long isoform	351	7E-95
		BAC03375	microtubule affinity-regulating kinase-like l	349	2E-94
		NP_113605	MAP/microtubule affinity-regulating kinase like 1, MARK4 serine/threonine protein kinase	349	2E-94
		AAD48007	serine/threonine protein kinase Kp78 splice variant CTAK75a	339	3E-91
		AAC33487	R31237_1, partial CDS	335	3E-90
	-	BAA96001	KIAA1477 protein	321	6E-86
		BAA07744	KIAA0096 gene product is related to a protein kinase.	285	5E-75
		P54646	S'-AMP-activated protein kinase, catalytic alpha-2 chain (AMPK alpha-2 chain).	285	6E-75
		AAF86944	HSNFRK	283	2E-74
NM_023499 Mm.214500 U:3.72 (YtoO) NP 075988.1		CAA75033	immunoglobulin lambda light chain	293	2E-78
		S25749	Ig lambda chain	276	2E-73
		BAC01857	immunoglobulin lambda light chain VLJ region	268	4E-71
	,				
-		BAC01837	immunoglobulin lambda light chain VLJ region	268	5E-71
		BAC01863	immunoglobulin lambda light chain VLJ region	268	SE-71
		BAC01842	immunoglobulin lambda light chain VLJ region	268	7E-71
		BAC01808	immunoglobulin lambda light chain VLJ region	265	6E-70
		BAC01859	immunoglobulin lanıbda light chain VLJ region	264	8E-70

2

0

ä

 \tilde{a}

		263		
	BAC01813	immunoglobulin lambda light chain VLJ region	263	2E-69
	BAC01822	immunoglobulin lambda light chain VLJ region	263	2E-69
	BAC01812.	immunoglobulin lambda light chain VLJ region	263	2E-69
	BAC01823	immunoglobulin lambda light chain VLJ region	263	2E-69
	BAC01834	immunoglobulin lambda light chain VLJ region	797	3E-69
	BAC01838	immunoglobulin lambda light chain VLJ region	797	3E-69
	BAC01820	immunoglobulin lambda light chain VLJ region	797	3E-69
	BAC01821	immunoglobulin lambda light chain VLJ region	797	3E-69
	BAC01840	immınoglobulin lambda light chain VLJ region	797	4E-69
	1203309A.	Ig lambda VI THO	197	5E-69
	BAC01824	immunoglobulin lambda light chain VLJ region	261	6E-69
	BAC01846	immunoglobulin lambda light chain VLJ region	197	6E-69
	BAC01841	immunoglobulin lambda light chain VLJ region	260	1E-68
	BAC01844	immunoglobulin lambda light chain VLJ region	260	1E-68
	BAC01862	immunoglobulin lambda light chain VLJ region	260	1E-68
	A42193	Ig lambda chain (BJP-DIA)	260	1E-68
	BAC01799	immunoglobulin lambda light chain VLJ region	260	1E-68
	BAC01860	immunoglobulin lambda light chain VLJ region	260	1E-68
	BAC01797	immunoglobulin lambda light chain VLJ region	259	2E-68
	BAC01814	immunoglobulin lambda light chain VLJ region	259	2E-68
,	BAC01810	immunoglobulin lambda light chain VLJ region	259	3E-68
	BAC01778	immunoglobulin lambda light chain VLJ region	259	3E-68
	BAC01792	immunoglobulin lambda light chain VLJ region	259	3E-68
	BAC01828	immunoglobulin lambda light chain VLJ region	259	3E-68
	BAC01847	immunoglobulin lambda light chain VLJ region	258	4E-68
	AAH33102	Similar to immunoglobulin lambda joining 3	258	4E-68
	BAC01854.	immunoglobulin lambda light chain VLJ region	258	5E-68
	BAC01839	irumunoglobulin lambda light chain VLJ region	258	5E-68

10 12 20

			7.10		
		BAC01855.	inmunoglobulin lambda light chain VLJ region	258	7E-68
		BAC01833.	immunoglobulin lambda light chain VLJ region	257	9E-68
		BAC01811.	inumunoglobulin lambda light chain VLJ region	257	1E-67
		S25744	Ig lambda chain	257	1E-67
		BAC01791	immunoglobulin lambda light chain VLJ region	257	1E-67
		S25738	Ig lambda chain	257	1E-67
-		BAC01819	immunoglobulin lambda light chain VLJ region	256	2E-67
	,	BAC01786	immunoglobulin lambda light chain VLJ region	256	2E-67
		BAC01806	immunoglobulin lambda light chain VLJ region	256	2E-67
		BAC01831	immunoglobulin lambda light chain VLJ region	256	2E-67
		BAC01818	immunoglobulin lambda light chain VLJ region	256	2E-67
-		BAC01779	immunoglobulin lambda light chain VLJ region	256	2E-67
		S25752	Ig lambda chain	256	2E-67
		1615309A	Ig lambda,anti-Rh(c).	256	2E-67
		BAC01845	immunoglobulin lambda light chain VLJ region	256	3E-67
		BAC01782	immunoglobulin lambda light chain VLJ region	256	3E-67
		CAA40943	ımmunoglobulin lambda light chain	256	3E-67
		BAC01830	immunoglobulin lambda light chain VLJ region	256	3E-67
		BAC01852	immunoglobulin lambda light chain VLJ region	256	3E-67
		S05270	Ig lambda chain precursor	256	3E-67
		BAC01856	immunoglobulin lambda light chain VLJ region	255	4E-67
		BAC01787	immunoglobulin lambda light chain VLJ region	255	4E-67
		AAH30983	Similar to immunoglobulin lambda joining 3	255	4E-67
		S25746	Ig lambda chain	255	4E-67
		BAC01789	immunoglobulin lambda light chain VLJ region	255	4E-67
		BAC01832	immunoglobulin lambda light chain VLJ region	255	4E-67
		BAC01800	immunoglobulin lambda light chain VLJ region	255	SE-67
		S25757	Ig lambda chain - human	254	6E-67
		BAC01777	immunoglobulin lambda light chain VLJ region	254	6E-67

IJ

 Ξ

7

C)

			2/1		
		BAC01796	immunoglobulin lambda light chain VLJ region	254	6E-67
		BAC01780	immunoglobulin lambda light chain VLJ region	254	6E-67
		BAC01835	immunoglobulin lambda light chain VLJ region	254	8E-67
		1	inmunoglobulin lambda light chain VLJ region	254	8E-67
		C01809	inmunoglobulin Jambda Jight chain VLJ region	254	8E-67
		S21066	Ig lambda chain V region	254	8E-67
		BAC01801	immunoglobulin lambda light chain VLJ region	254	8E-67
		1LILA	Chain A, Bence Jones Protein Cle, A Lambda Iii Immunoglobulin Light-Chain Dimer.	254	8E-67
		BAC01853	immunoglobulin lambda light chain VLJ region	254	8E-67
			I		
NM_009255 Mm.3093	U:3.6 (5to19)	XP_059422.1	similar to tropomyosin, fibroblast - human	169	0
NP 033281.1					
		P07093	Glia derived nexin precursor (GDN) (Protease nexin I) (PN-1) (Protease inhibitor 7)	684	0
		A26061	glia-derived neurite promoting factor precursor	682	0
		pdb/1DB2	Plasminogen activator inhibitor-1 precursor (PAI-1) (Endothelial plasminogen activator inhibitor)	310	4E-83
			(PAI)		
		CAA28444.1	plasminogen activator inhibitor	310	6E-83
		pdb 11.35	Chain A, 1.8a Resolution Structure Of Latent Plasminogen Activator Inhibitor-1(Pai-1)	310	6E-83
		NP_000593.1	000593.1 serine (or cysteine) proteinase inhibitor, clade E (nexin, plasminogen activator inhibitor type 1),	310	6E-83
			member 1; plasminogen activator inhibitor, type I		·
		AAA60008.1	A60008.1 prebeta-migrating plasminogen activator inhibitor	310	6E-83
		AAA60009.1	AAA60009.1 plasminogen activator inhibitor 1	308	1E-82
		pdb 9PAI	Plasminogen activator inhibitor-1 precursor (PAL-1) (Endothelial plasminogen activator inhibitor)	308	2E-82
			(PAI)		
		pdb IA7C	Plastumogenactivator inhibitor-1 precursor (PAI-1) (Endothelial plasminogen activator inhibitor)	308	2E-82
			(PAI)	_ {	
-		pdb 1B3K	Plasminogen activator inhibitor-1 precursor (PAL-1) (Endothelial plasminogen activator inhibitor)	307	3E-82
			(PAI)		
		pdb 1DVM	Chain A, Active Form Of Human Pai-1	305	2E-81

2

10

5

 \sim

		CAA31208.1	A31208.1 PAI precursor polypeptide	305	2E-81
		NP_005016.1	005016.1 ne (or cysteine) proteinase inhibitor, clade I (neuroserpin), member 1; protease inhibitor 12	243	7E-63
			(neuroserpin)		
		AAH18043.1	AAH18043.1 serine (or cysteine) proteinase inhibitor, clade I (neuroserpin), member 1	240	5E-62
		NP_006208.1	006208.1 protease inhibitor 14; pancpin	219	1E-55
NM_021468 Mm.42188	U:3.58 (MtoO)	NP_006368	UNC13 (C. elegans)-like; homolog of rat Munc13 (diacylglycerol-binding)	2958	0
NP 067443.1					
		BAA82984	KIAA1032 protein	1598	0
		XP_038604	similar to KIAA1032 protein	1478	0
		303675	unnamed protein product	1404	0
		XP_085234	similar to Munc13-3 protein - rat	1328	0
·		CAD39069	hypothetical protein	915	0
NM_007643 Mm.18628 NP 031669.1	U:3.57 (YtoO)	P16671	Platelet glycoprotein IV (GPIV) (GPIIB) (CD36 antigen) (PAS IV) (PAS-4 protein)	798	0
		NP_000063.1	.000063.1 CD36 antigen (collagen type I receptor, thrombospondin receptor); CD36 antigen (collagen type	196	0
			Γ		
		159613	cell adhesion receptor CD36	791	0
		AAM14636.1	AAM14636.1 CD36 antigen (collagen type I receptor, thrombospondin receptor)	780	0
		NP_005497.1	005497.1 scavenger receptor class B, member 2; CD36 antigen (collagen type I receptor, thrombospondin	271	3E-72
•			receptor) -; CD36 antigen (collagen type I receptor, thrombospondin receptor)-like 2 (lysosomal	-	
			integral membrane protein II)		
		A56525	lysosomal integral membrane protein II	271	3E-72
		NP_005496.2	.005496.2 scavenger receptor class B, member 1; CD36 antigen-like 1; scavenger receptor class B type 1;	255	2E-67
		·	CD36 antigen (collagen type I receptor, thrombospondin receptor)-like 1		
-		A48528	membrane glycoprotein CLA-1 protein long form precursor	252	2E-66
A K 0 0 7 2 9 3 Mn. 159753 U:3.56 (5to11)	3 U:3.56 (5to11)	BAB67772.1	BAB67772.1 KIAA1879 protein	189	8E-47
BAB24937.1					

Ŋ

10

5

 $\vec{\sim}$

	3E-70	3E-40	1,	4E-40	0			0	0	0	0	0	0	1E-143	1E-143	1E-109	7E-85	7E-85	3E-78	1E-70
	766	165	32	<u>a</u>	1875		1875	833	728	726	969	969	889	510	510	398	317	317	295	270
273	Similar to RIKEN cDNA 2010001M09 gene	NP_000598.1 orosomucoid 1 precursor; Orosomucoid-1 (alpha-1-acid glycoprotein-1); alpha-1-acid		Orosomucona 1	NP_055370.1 transient receptor potential cation channel, subfamily M, member 5; MLSN1 and TRP-related;	MLSN1- and TRP-related	CAB66342.1 LTRPC5 protein	NP_060106.2 transient receptor potential cation channel, subfamily M, member 4	AL02142.1 TRP-related cation influx channel	BAA90907.1 unnamed protein product	BAA95563.1 transient receptor potential-related channel 7, a novel putative Ca2+ channel protein	NP_003298.1 transient receptor potential cation channel, subfamily M, member 2; transient receptor potential-related channel 7, a novel putative Ca2+ channel protein; transient receptor potential channel 7	CAD01139.1 putative TRP cation channel	LTRPC6	transient receptor potential cation channel, subfamily M, member 8	NP_060132.3 transient receptor potential cation channel, subfamily M, member 6	channel-kinase 1	XP_030709.6 similar to LTRPC7	AB15429.1 unnamed protein product	melastatin 1
	AAH21275	NP_000598.1	4 A LL76730	AAH 20238.1	NP_055370.1		CAB66342.1	NP_060106.2	AAL02142.	BAA90907.	BAA95563.	NP_003298.	CAD01139.	BAB86335.1 LTRPC6	NP_076985.	NP_060132.	AAK19738.	XP_030709.	BAB15429.	AAC80000.1 melastatin 1
	U:3.37 (YtoO)	U:3.35 (7to19)		- 1	NM_020277 Mm. 143747 U:3.35 (5to11)															
	Mm.27252	Mm.57239			Mm. 14374															
	A K 0 0 8 0 1 6 Mm.27252 AK008016	NM_013623 Mm.57239	NP 038651.1		NM_020277	NP 064673.1								·						

27.

			7.7		
		NP_002411.2	002411.2 transient receptor potential cation channel, subfamily M, member 1, melastatin 1	270	1E-70
NM_007809 Mm.1262	U:3.27 (YtoO)	A26366	steroid 17alpha-monooxygenase (EC 1.14.99.9) cytochrome P450 17	710	0
NP 031835.1					
		AAA36405	cytochrome P450c17	709	0
		AAA52140	steroid 17-alpha-hydroxylase	90/	0
		CAA26458	cytochrome P(1)-450	248	2E-64
		O4HU6	aryl hydrocarbon (benzo[a]pyrene) hydroxylase (EC 1.14.14) cytochrome P450 1A1	248	3E-64
		AAA52139	cytochrome P-450-1	246	1E-63
J00544 Mm.1192	U:3.24 (YtoO)	NP_653247	inumunoglobulin J polypeptide, linker protein for immunoglobulin alpha and mu polypeptides	245	8E-64
AAA38673.1					
		AAA58902	Ig J-chain	228	1E-58
NM_031494 Mm.26595	U:3.21 (YtoM)	Q9NSD4	Z275_HUMAN Zinc finger protein 275	840	0
NP_113682.1		-			
		BAC04309.1	unnamed protein product	363	2E-98
		P52742	Z135_HUMAN Zinc finger protein 135	360	2E-97
		CAD39111.1	hypothetical protein	357	2E-96
	. 1	AAF75235.1	AF244088_1 zinc finger protein	357	2E-96
		AAH10996.1	AAH10996 zinc finger protein 16 (KOX 9)	357	2E-96
		AAC50252.1	zinc finger protein ZNF132	357	2E-96
		AAH28377.1	AAH28377.1 hypothetical protein FLJ14855	349	4E-94
		AAH05868.1	AAH05868.1 Similar to zinc finger protein 304	348	SE-94
		NP_694563.1	694563.1 hypothetical protein FLJ30726	347	2E-93
		NP_653290.2	653290.2 hypothetical protein FLJ32191	346	2E-93
		AAH32863.1	Unknown (protein for MGC:33794)	346	2E-93
		BAC11133.1	BAC11133.1 unnamed protein product	343	2E-92
		NP_660355.1	660355.1 Zinc finger protein 93 (Zinc finger protein HTF34)	343	2E-92
		XP_209899.1	209899.1 similar to Zinc finger protein 184	343	3E-92

10 10 20 20

Ŋ

	275		
AAA97578.1	ZNF157	342	3E-92
BAB71272.1	umamed protein product	342	4E-92
NP_689817.1	NP_689817.1 hypothetical protein FLJ35863	342	4E-92
AAC34327.1	F23269_2	342	4E-92
CAC88160.1 b	bB479F17.1 (zinc finger protein 1S7 (HZF22))	342	4E-92
XP_209140.1	209140.1 similar to F23269_2	342	4E-92
AAH42500.1	Similar to zinc finger protein 43 (HTF6)	342	4E-92
AAH09433.1	zinc finger protein 331	341	7E-92
AAF78075.1	KRAB zinc finger protein	341	7E-92
NP_061121.1	zinc finger protein ZFP	340	1E-91
BAB71257.1	unnamed protein product	340	1E-91
AAH45649.1	Similar to hypothetical protein FLJ32191	340	1E-91
CAC06610.1	zinc finger protein 304	340	2E-91
BAB21801.1	KLAA1710 protein	339	3E-91
AAH43151.1	Similar to zinc finger protein 208	339	3E-91
Q9C0F3	Hypothetical zinc finger protein KIAA1710	339	3E-91
XP_209718.1	similar to zinc finger protein 184 (Kruppel-like)	338	6E-91
BAC04418.1	unnamed protein product	338	6E-91
AAH28136.1	AAH28136.1 Similar to hypothetical protein MGC10520	338	8E-91
AAH04480.1	Unknown (protein for MGC:10520)	338	8E-91
XP_087503.1	XP_087503.1 similar to zinc finger protein 91 (HPF7, HTF10)	338	8E-91
BAC04610.1	unnamed protein product	337	1E-90
XP_086070.1	XP_086070.1 similar to Zinc finger protein similar to Zinc finger protein 93 (Zinc finger protein HTF34)	337	1E-90
P35789	ZN93_HUMAN Zinc finger protein 93	337	1E-90
NP_085116.1	085116.1 hypothetical protein FLJ21628	337	2E-90
XP_065116.3	similar to zinc finger protein 91 (HPF7, HTF10)	.337	2E-90
AAH33849.1	similar to Zinc finger protein 268 (Zinc finger protein HZF3)	336	3E-90
BAB70771.1	unnamed protein product	336	3E-90
CAD38551,I	CAD38551,1 hypothetical protein	335	4E-90

		2/2		~
/V	AK52068.1	AK52068.1 ZNFPHEX133 protein	335	4E-90
B,	AB15582.1		335	4E-90
X	P_032054.2	similar to zinc finger protein	335	5E-90
B	AB85542.1	KIAA1956 protein	335	5E-90
A	AD23607.1	AC007228_2 BC37295_	335	5E-90
0	05481	ZN91_HUMAN Zinc finger protein 91 (Zinc	335	5E-90
×	P_085836.1	similar to Hypothetical zinc f	335	5E-90
. B.	AB71096.1	umamed protein product	335	5E-90
A	AH08297.1	Similar to hypothetical protein	335	7E-90
B	AA92634.1	KIAA1396 protein	335	7E-90
× .	P_030378.2	similar to zinc finger protein 28; zinc finger factor X6	335	7E-90
B	AB47458.1	BAB47458.1 KIAA1829 protein	335	7E-90
V	AD39268.1	BC331191_1	334	9E-90
Z	P_689690.1	hypothetical protein FLJ36991	334	9E-90
Z	IP_055713.1	KIAA0961 protein	334	9E-90
Z	IP_055295.1	zinc finger protein AF020591	334	9E-90
X	P_209968.1	similar to DKFZP572C163 protein	334	1E-89
X	P_091895.5	similar to KIAA1947 protein	334	1E-89
2	JP_150630.1	KRAB zinc finger protein KR18	334	1E-89
B	AB13437.1	KIAA1611 protein	334	1E-89
B	AB85533.1	KIAA1947 protein	334	1E-89
B	AB15732.1	FLJ00032 protein	334	1E-89
)	AD38678.1	hypothetical protein	333	2E-89
)	AD36956.1	zinc finger protein 33b	333	2E-89
)	AB45722.1	hypothetical protein	333	2E-89
B	3AB14911.1	unnamed protein product	333	2E-89
I	114757	hypothetical protein DKFZp572C163.1	333	3E-89
<u> </u>	NP_666016.1	zinc finger protein 23; zinc finger protein 32; zinc finger protein 359	333	3E-89
B	3AB14145.1	AB14145.1 unnamed protein product	333	3E-89

Ŋ

-

5

20

Ŋ

			277		
NM_008161 Mm.7156	U:3.13 (YtoO)	BAA00525.1	BAA00525.1 glutathione peroxidase	397	1E-109
NP 032187.2					
	٠	JQ0476	glutathione peroxidase (EC 1.11.1.9) 3, precursor	397	1E-109
	, 12	AAF43005.1	extracellular glutathione peroxidase	390	1E-107
		CAA06463.1 g	glutathione peroxidase type 5 (GPX5)	301	1E-80
		BAA03864.1	plasma glutatlione peroxidase	281	1E-74
		XP_167146.1	XP_167146.1 similar to EPIDIDYMAL SECRETORY GLUTATHIONE PEROXIDASE PRECURSOR	202	8E-51
			(EPIDIDYMIS-SPECIFIC GLUTATHIONE PEROXIDASE-LIKE PROTEIN) (EGLP)		
NM_025724 Mm.56430	U:3.12 (MtoO)	NM_013559	NM_013559 Unknown (protein for MGC:26598)	262	2E-68
NP 080000.1					
		AAH24183	Similar to RIKEN cDNA 4921510H08 gene	262	2E-68
NM_011125 Mm6105	U:3.1 (YtoO)	AAH19847.1	AAH19847.1 phospholipid transfer protein	744	0
NP 035255.1					
		NP_006218.1	NP_06218.1 phospholipid transfer protein	744	0
		CAC36020.1	CAC36020.1 dJ337O18.1.2 (Phospholipid Transfer Protein (Lipid Transfer Protein II) (isoform 2))	634	0
		AAH05045.1	AAH05045.1 Similar to phospholipid transfer protein	633	0
NM_012006Mm.1978	U:3.07 (5to7)	XP_170752.1	similar to peroxisomal long-chain acyl-coA thioesterase; peroxisomal long-chain acyl-coA	602	1E-172
NP 036136.1			thioesterase; putative protein		
		P49753	Peroxisomal acyl-coenzyme A thioester hydrolase 2a (Peroxisomal long-chain acyl-coA	009	1E-171
-			thioesterase 2) (ZAP128)		
		AAH06500.1	AAH06500.1 Unknown (protein for MGC:2366)	009	1E-171
		NP_006812.2	NP_006812.2 peroxisonnal long-chain acyl-coA thioesterase; peroxisonnal long-chain acyl-coA thioesterase;	599	1E-171
			putative protein		
		BAA91989.1	BAA91989.1 unnamed protein product	865	1E-171
	Janes				

			2.10		
		NP_689544.1	689544.1 hypothetical protein FLJ31235	494	1E-139
		AAC42007.1	242007.1 ORF; putative	405	1E-113
		XP_090885.1	record was removed	280	4E-75
NM_008361 Mm.22150	U:3.05 (5to7)	NP_000567.1	000567.1 interleukin 1, beta proprotein; preinterleukin 1 beta; interleukin 1; catabolin	352	3E-97
NP 032387.1					
		P01584	interleukin 1, beta proprotein; preinterleukin 1 beta; interleukin 1; catabolin	350	1E-96
		AAA59136.1 interleukin	interleukin 1	345	6E-95
				3	i c
		AAC03536.1	interleukin 1 beta	240	3E-63
		1827779	Chain , Interleukin-1 Beta From Joint X-Ray And Nmr Refinement	239	3E-63
		230947	Chain, Interleukin-1Beta (IL-1Beta) (Mutant With Cys 8 Replaced By Ala (C8A)	239	3E-63
		494152	Chain , Interleukin-1 Beta (Human) Mutant With Thr 9 Replaced By Gly (T9g)	239	3E-63
		230410	Chain, Interleukin-1Beta (IL-1Beta) (Mutant With Cys 71 Replaced By Ala) (C71A)	236	3E-62
		230798	Chain, Interleukin-1Beta (IL-1Beta) (Mutant With Cys 71 Replaced By Ser) (C71S)	236	4E-62
NM_013559 Mm.34828	U:2.97 (YtoO)	AAC18044	antigen NY-CO-25	1529	0
NP 038587.1					
		BAA13192	similar to mouse heat shock protein 105 kDa beta	1524	0
		NP_006635	heat shock 105kD; heat shock 105kD alpha; heat shock 105kD beta	1465	0
	•	XP_114482	similar to HEAT SHOCK 70 KDA PROTEIN 4 (HEAT SHOCK 70-RELATED PROTEIN APG-	1025	0
			2) (HSP70RY)		
		BAA75062	apg-2	1021	0
		BAA75063	apg-1	951	0
		AAH40560	heat shock protein (hsp110 family)	948	0
		156208	heat shock protein 70	905	0
		CAA47886	HS24/P52	674	0
		CAA47885	HS24/P52	395	1E-108
AF127033 Mm.3760	U:2.97 (YtoO)	NP_004095	fatty acid synthase	3961	0
AAG02285.1					

Ŋ

2

 $\tilde{\mathbf{z}}$

WO 2005/000335 PCT/US2004/017322

		!	0/10		
		G01880	fatty-acid synthase (EC 2.3.1.85) (version 2)	3945	0.
*		A57788	enoyl-[acyl-carrier-protein] reductase (NADPH2, B-specific) (EC 1.3.1.10) (version 1)	3778	0
		AAH07267	Unknown (protein for IMAGE:3138929)	1533	0
		AAB35516	fatty acid synthase; FAS	728	0
		AAH07305	Unknown (protein for MGC:15706)	685	0
		AAH14634	Unknown (protein for IMAGE:353581)	553	1E-155
NM_010062 Mm.41853	U:2.89 (5to11)	NP_001366.1	NP_001366.1 deoxyribonuclease II, lysosomal; DNase II, lysosomal	520	1E-147
NP 034192.1					
		T45071	hypothetical protein R31240_2 [imported]	494	1E-139
		NP_067056.1	deoxyribonuclease II beta, isoform 1 precursor; DNase II-like acid DNase; endonuclease DLAD	227	5E-59
		AAL34449.1	endonuclease DLAD	227	SE-59
NM_011704 Mm.27154	U:2.87 (5to7)	NP_004657.1	NP_004657.1 vanin 1 precursor; Vannin 1; pantetheinase	795	0
NP 035834.1					
		AAF21453.1	AAF21453.1 Tiff66	793	0
	1	CAB40075.1	dJ55C23.1 (vanin 1)	793	0
					,
		NP_060869.1	NP_060869.1 vanin 3 isoform 1 precursor; VNN3 protein; pantetheinase	657	0
		CAB40076.1	CAB40076.1 dJ55C23.2 (vanin 2)-	639	0
		NP_004656.2	NP_004656.2 vanin 2, isoform 1 precursor; Vannin 2;	638	0
		CAA10569.1	VNN2 protein	635	0
		NP_511043.1	NP_511043.1 vanin 2, isoform 2; Vannin 2;	597	1E-169
		P43251	Biotinidase precursor	382	1E-105
		NP_000051.1	NP_000051.1 biotinidase precursor	382	1E-105
		CAC33872.1	dJ55C23.5.1 (vanin 3, isoform 1)	342	1E-92
A K 0 1 8 6 9 5 Mm. 29805	U:2.85 (YtoM)	Q14156	Hypothetical protein KIAA0143	461	0
	•				

5

		280		
NP_598527.1				
	XP_035825	similar to Hypothetical protein KIAA0143	461	٦
	BAA76797	KJAA0953 protein	266	1E-136
		similar to KIAA0953 protein	792	1E-136
NM 011674 Mm.5160 U:2.84 (7to19)	9) Q16880	2-hydroxyacylsphingosine 1-beta-galactosyltransferase precursor(UDP-galactose-ceramide	1042	0
NP 035804 1		galactosyltransferase) (Ceramide UDP-galactosyltransferase) (Cerebroside synthase).		
	NP_003351.1		1039	0
	JC5423	2-hydroxyacylsphingosine 1-beta-galactosyltransferase (EC 2.4.1.45)	1031	0
	075795	UDP-glucuronosyltransferase 2B17 precursor, microsomal (UDPGT) (C19-steroid specific UDP-	332	1E-89
	•	glucuronosyltransferase).		
	AAC50077.1	C50077.1 UDP glucuronosyltransferase precursor	329	9E-89
	AF180322_1	UDP-glucuronosyltransferase 2B15	329	9E-89
	AF297093 4	AF297093_4 UDP glucuronosyltransferase 1A7	326	8E-88
	NP 066307.1	066307.1 UDP glycosyltransferase 1 family, polypeptide A9	323	SE-87
	AAB81536.1	UDP-glucuronosyltransferase 1A7	323	5E-87
	S11309	glucuronosyltransferase (EC 2.4.1.17)	321	3E-86
	JC5656	UDP glucuronosyltransferase (EC 2.4.1) 1A10 precursor	319	7E-86
	AF462267_1	UDP-glucuronosyltransferase UGT1A8*2	319	1E-85
	AF297093_1	UDP glucuronosyltransferase 1A8	319	1E-85
	AAH20971	Similar to UDP glycosyltransferase 1 family, polypeptide A9	318	1E-85
	AAB84259.1	UDP-glucuronosyltransferase 1A8	316	8E-85
	NP_006789.	006789.1 UDP glycosyltransferase 2 family, polypeptide A1; UDP glucuronosyltransferase 2 family,	, 315	1E-84
		polypeptide A1		1
	P36509	UDP-glucuronosyltransferase 1-2 precursor, microsomal (UDPGT)(UGT-1B) (UGT1*2) (UGT1-	315	1E-84
		(22) (UGT1.2) (UGT1A2) (UGT1B) (HLUGP4).		
	P16662	UDP-glucuronosyltransferase 2B7 precursor, microsomal (UDPGT)(3,4-catechol estrogen	n 313	4E-84
		specific) (UDPGTH-2).		
	AAB81537.	AB81537.1 UDP-glucuronosyltransferase 1A10	313	5E-84

. 25

S

		281		
	AAH30974.1	AH30974.1 UDP glycosyltransferase 2 family, polypeptide B7	313	7E-84
	AAB19791.2	.B19791.2 UDP-glucuronosyltransferase	312	9E-84
	S17512	glucuronosyltransferase (EC 2.4.1.17)	311	3E-83
	AAF78145.1	UDP-glucuronosyltransferase	310	6E-83
	AAC95002.1	UDP-glucuronosyltransferase 2B4 precursor	309	8E-83
	9190Nt	glucuronosyltransferase (EC 2.4.1.17) 2B-4 precursor	309	1E-82
	JN0620	JN0620 UDP-glucuronosylfransferase (EC 2.4.1) 2B-10 precursor	308	2E-82
	AAC32272.1	AAC32272,1 UDP glucuronosyltransferase 2B4 precursor	306	8E-82
	AF297093_5	AF297093_5 UDP glucuronosyltransferase 1A6	305	1E-81
	AAC27891.1	UDP-glucuronosyltransferase 2B	305	2E-81
	JE0200	orphan UDP-glucuronosyltransferase (EC 2.4)	304	2E-81
	AF297093_6	AF297093_6 UDP glucuronosyltransferase 1A5	304	3E-81
	AAH19861	Unknown (protein for MGC:29860)	304	3E-81
	BAB15179.1	unnamed protein product	304	3E-81
NM_023455 Mm.154782 U:2.75 (51019)	NP_057431.1	NP_057431.1 putative N-acetyltransferase Camello 2	223	4E-58
NF 0/3944.1	NP 003951.2	N-acetyltransferase 8; kidney- and liver-specific gene product; kidney- and liver-specific gene	216	3E-56
	BAA71643.1	BAA71643.1 GLA	216	4E-56
	AAH12626.1	kidney- and liver-specific gene	214	1E-55
	T44342	hypothetical protein TSC501 [imported]	214	1E-55
NM_023478 Mm.46214 U:2.74 (5to19)	075631	Uroplakin III precursor (UPIII)	496	1E-140
NP 075967.1				
	NP_008884.1 uroplakin 3	uroplakin 3	496	1E-140
	BAÁ25678.1	uroplakin 3	439	1E-123
NM_016774 Mm.103838 U:2.73 (YtoM)	P06576	ATP synthase beta chain, mitochondrial precursor	893	0
NP 058054.1				

		779100 an	ATD centhace It transconting mitrohondrial El complex betanchmanide. ATD centhace It	008	
		INI_0010011	A LE SYMMASE, 117 d'ambroumB, minochomman et complex, octa porypépique, A LE Symmase, 117		>
	-		transporting, mitochondrial F1 complex, beta		
		AAA51808	ATP synthase beta subunit	879	0
	1	CAA29095	beta-subunit (AA 1-312)	265	1E-170
NM_011146 Mm.3020	U:2.68 (5to11)	NP_056953.2	P_056953.2 peroxisome proliferative activated receptor gamma, isoform 2; PPAR-gamma; peroxisome	953	0
NP 035276.1			proliferator activated receptor gamma		
		BAA18949.1	BAA18949.1 PPAR garuma2	626	0
		S42489	peroxisome proliferator activated receptor	922	0
		CAA62152.1	peroxisome proliferator activated receptor gamma	916	0
		NP_005028.3	NP_005028.3 peroxisome proliferative activated receptor gamma, isoform 1; PPAR-gamma; peroxisome	914	0,
			proliferator activated receptor gamma		
		BAA23354.1	BAA23354.1 peroxisome proliferator activated-receptor gamma	904	0
		20150106	Chain A, Crystal Structure Of The Ligand Binding Domain Of Human Ppar-Gamma In Complex	511	1E-144
			With The Agonist Az 242		
NM_013771 Mm.23335	U:2.67 (YtoM)	CAB51858	ATP-dependent metalloprotease YMB1L	1341	0
NP 038799.1					
		NP_647473	YME1-like 1 isoform 1; ATP-dependent metalloprotease FtsH1 homolog	1286	0
		AAK57555	ATP-dependent metalloprotease FtsH1 homolog	1285	0
		AAH07795	Similar to YME1-like 1 (S. cerevisiae)	1224	0
		AAD20962	FtsH homolog	992	0
		CAB99462	putative ATPases	166	0
-		CAC19650	bA145E8.2 (YME1 (S.cerevisiae)-like 1)	842	0
A K 0 0 2 9 7 9 Mm. 195881 U:2.67 (5to19)	31 U:2.67 (5to 19)	NP_056537.1 calcyon	calcyon	336	5E-92
BAB22492.1					
A K 0 0 5 6 0 9 Mm. 45109	U:2.62 (5to19)	XP_059692	similar to RIKEN cDNA 1700001L19 [Mus musculus]	228	9E-59
BAB24148.1					

Ŋ

0

20

				283		
X03796		U:2.61 (YtoM)	AAC09348	aldolase C	443	1E-123
CAA27422.1						-
			CAA30270	aldolase C	443	1E-123
			CAA28861.1	aldolase A (AA 1-364)	388	1E-107
			1ALD	Aldolase A (E.C.4.1.2.13)	386	1E-106
		1	CAA30979	aldolase A	380	1E-104
			AAB59377	aldolase B	335	4E-91
			ADHUB	fructose-bisphosphate aldolase (EC 4.1.2.13) B	335	4E-91
	,		AAA51691	aldolase B	334	9E-91
			BAA00125	aldolase B	334	9E-91
			10050	Chain Q, Fructose 1,6-Bisphosphate Aldolase From Human Liver	333	1E-90
			AAH29399	Similar to aldolase B, fructose-bisphosphate	333	1E-90
			1313294A	aldolase B	328	4E-89
NM_019415 Mm.182905 U.2.6 (5to11)	Mm.182905	U:2.6 (5to11)	P55017	Solute carrier family 12 member 3 (Thiazide-sensitive sodium-chloride cotransporter) (Na-Cl	1795	°
NP 062288.1				symporter)		i
			G01202	NaCl electroneutral Thazide-sensitive cotransporter	1792	0
			NP_000330.1	000330.1 solute carrier family 12 (sodium/chloride transporters), member 3; Solute carrier family 12	1792	0
				(sodium/potassium/chloride transporters),		
·			NP_001037.1	001037.1 solute carrier family 12 (sodium/potassium/chloride transporters), member 2; Solute carrier family	1023	0
				12 (sodium/potassium/chloride transporters),		
	-		NP_000329.1	000329.1 sodium potassium chloride cotransporter 2; Solute carrier family 12 (sodium/potassium/chloride 1022	1022	°
				transporters),		
			AAH33003.1	AAH33003.1 Similar to solute carrier family 12 (sodium/potassium/chloride transporters), member 2	944	0
			PC4180	thiazide-sensitive sodium-chloride cotransporter - human (fragment)	999	°
			NP_006589.1	006589.1 solute carrier family 12 (potassium/chloride transporters), member 7; potassium/chloride	316	1E-84
				transporter KCC4		

٣U

С

72

S

	-	AAL32454.1	.32454. I sodium-potassium-chloride cotransporter	296	1E-78
		BAA86490.1	186490.1 KIAA1176 protein	271	4E-71
		AAG43493.1	343493.1 electroneutral potassium-chloride cotransporter KCC2	271	4E-71
		NP_065759.1	065759.1 solute carrier family 12, (potassium-chloride transporter) member 5	271	4E-71
A K 0 0 9 9 3 7 Mm 196576 U:2.57 (YtoM) RAB26596 1	76 U:2.57 (YtoM)	AAH07220	hypothetical protein FLJ12118	857	0
		NP_078813	hypothetical protein FLJ12118	853	0
		BAB93499	OK/SW-CL.10	664	0
		NP_001742	cysteine-tRNA ligase, isoform b; cysteine translase; cysteine-tRNA synthetase	233	6E-61
		NP_644802	cysteine-tRNA ligase, isoform a; cysteine translase; cysteine-tRNA synthetase	233	6E-61
		AAA73901	cysteinyl-tRNA synthetase.	227	4E-59
NM_023137 Mm.140210 U.2.56 (YtoO)	10 U:2.56 (YtoO)	AAD52982	ubiquitin-like protein FAT10	211	1E-53
NP 075626.1					
		AAH12472	Sinular to diubiquitin	210	2E-53
A K 0 1 5 7 5 0 Mm.89655	5 U:2.56 (YtoO)	1HY3A	Chain A, Crystal Structure Of Human Estrogen Sulfotransferase V269e Mutant In The Presence	497	1E-139
BAB29956.1			OfPaps		
		JC2229	estrogen sulfotransferase (EC 2.8.2)	494	1E-138
		AAH27956.1	sulfotransferase, estrogen-preferring	492	1E-138
		AAB65154.1	thyroid hormone sulfotransferase	323	4E-87
		BAA24547.1	ST1B2	323	4E-87
		AAH10895.1	Unknown (protein for MGC:13356)	322	1E-86
		AAA67895.1	phenol sulfotransferase	315	1E-84
	-	P50225	Phenol-sulfating phenol sulfotransferase 1 (P-PST) (Thermostable phenol sulfotransferase) (Ts-	313	3E-84
			PST) (HAST1/HAST2) (ST1A3).		
		AAB31316.1	aryl sulfotransferase ST1A2	313	5E-84
		AAC50480.1	phenol sulfotransferase	313	5E-84
		AAH00923.1	AAH00923.1 sulfotransferase family, cytosolic, 1A, phenol-preferring, member 1	312	6E-84

WO 2005/000335 PCT/US2004/017322

2E-506E-84 1E-83 3E-83 6E-83 1E-82 2E-82 2E-82 2E-80 9E-80 7E-51 7E-51 4E-71 7E-51 2E-7 2E-7 9E-7 308 300 298 202 311 309 307 307 290 290 288 202 202 201 Chain A, Crystal Structure Of The Human Hydroxysteroid Sulfottansferase In The Presence Of XP_065757.1 similar to sulfotransferase, phenol preferring 2; Phenol sulfotransferase 1c1 [Rattus norvegicus] SULTIC sulfotransferase; sulfotransferase family, cytosolic, 1C, member C2 hydroxysteroid sulfotransferase SULT2B1b hydroxysteroid sulfotransferase SULT2B1a AAB18753.1 phenol-preferring phenol sulfotransferase 2 hydroxysteroid sulfotransferase SULT2B1a CAA59147.1 phenol-sufating phenosulfotransferase aryl sulfotransferase (EC 2.8.2.1) AAB31317.1 aryl sulfotransferase ST1A3 BAB93491.1 sulfotransferase family 1A arylamine sulfotransferase AAB09758.1 phenol sulfotransferase aryl sulfotransferase sulfotransferase 1C2 sulfotransferase 1C1 sulfotransferase AAC51149.1 NP 006579.1 AAF72804.1 AAC78499.1 AAC00409.1 AAC78498.1 AAC99987.1 AAC78553.1 AAF72810.1 A55451 IEFHA

Ŋ

2E-50 2E-50 2E-50 IE-145 1E-83 2E-50 1E-83 516 201 201 201 201 NP_003158.1 sulfotransferase family, cytosolic, 2A, dehydroepiandrosterone(DHEA) -preferring, member 1; Chain A, Crystal Structure Of Human Dehydroepiandrosterone Sulfotransferase In Complex With sulfotransferase family 2A, dehydroepiandrosterone (DHEA) -preferring, member 1 diacylglycerol O-acyltransferase 2 like 1; iacylglycerolacyltransferase 2-like AAB23169.2 | alcohol/hydroxysteroid sulfotransferase; hSTa dehydroepiandrosterone sulfotransferase diacylglycerol acyltransferase 2 hypothetical protein Substrate. AAA75491.1 AAK84176. NP_477513 CAD38961 1J99A U:2.55 (YtoO) A K 0 0 2 6 9 3 Mm.41325 BAB22288.

25

10 10 20

	-	CAD13492	bA351K23.5 (novel protein)	258	1E-67
		BAB15436	unnamed protein product	241	1E-62
		AAD45832	similar to predicted proteins AAB54240 ID:g2088822) and S67138 (PID:g2132925)	208	1E-52
NM_011214 Mm.4860	U:2.54 (MtoO)	JC5290	protein-tyrosine-phosphatase (EC 3.1.3.48)	2649	0
NP 035344.1					
-		AAB07074.1	receptor protein tyrosine phosphatase psi	2647	0
		NP_573439.1	NP_573439.1 protein tyrosine phosphatase, receptor type, U isoform 1 precursor; protein tyrosine phosphatase	2642	0
			I; protein tyrosine phosphatase receptor omicron		
		NP_573438.1	573438.1 protein tyrosine phosphatase, receptor type, U isoform 2 precursor; protein tyrosine phosphatase	2635	0
			J; protein tyrosine phosphatase receptor omicron		
		NP_005695.2	005695.2 protein tyrosine phosphatase, receptor type, U isoform 3 precursor; protein tyrosine phosphatase	2634	0
			J; protein tyrosine phosphatase receptor omicron		
		AAC51938.1	protein tyrosine phosphatase receptor omicron	2618	0
		CAA65832.1	receptor phosphatase PCP-2	2505	0
		CAA65016.1 FMI protein	FMI protein .	2437	0
		NP_002835.2	NP_002835.2 protein tyrosine phosphatase, receptor type, K precursor; protein-tyrosine phosphatase, receptor	r 1667	0.
			type, kappa; protein-tyrosine phosphatase kappa; protein-tyrosine phosphatase kappa		
			precursor		
		Q15262	Protein-tyrosine phosphatase kappa precursor (R-PTP-kappa).	1664	0
		AAC37599.1	protein tyrosine phosphatase	1662	0
		JC6312	protein-tyrosine-phosphatase (EC 3.1.3.48)	1645	0
		CAB51346.1	dJ437116.1.1 (Protein-tyrosine phosphatase (isoform 1))	1488	0
		CAB51348.1	dJ437I16.1.3 (Protein-tyrosine phosphatase (isoform 3))	1480	0
		CAB51347.1		1480	0
		S17669	protein-tyrosine-phosphatase (EC 3.1.3.48), receptor type mu precursor	1478	0
		NP_002836.2	_002836.2 protein tyrosine phosphatase, receptor type, M precursor; protein tyrosine phosphatase, receptor	r 1478	0
			type, mu polypeptide; protein tyrosine phosphatase mu precursor		
		1717216B	receptor-like Tyr phosphatase	1475	0

10

			787		
	BA	BAA22952.2 KIAA0283	KIAA0283	1442	0
,	dN.	008981.2	008981.2 protein tyrosine phosphatase, receptor type, T, isoform 2 precursor; receptor protein tyrosine	1435	0
			phosphatase		
	ĄĄ	AD09421.2	receptor protein tyrosine phosphatase	1427	0
	CA	\D54117.1	CAD54117.1 dJ480J14.2.1 (protein tyrosine phosphatase, receptor type, K(R-PTP-KAPPA, protein tyrosine	706	0
			phosphatase kappa, protein tyrosine phosphatase kappa precursor), variant 1)		
	CA	A19666.1	dJ707K17.1 (Protein tyrosine phosphatase, receptor type, T(RPTPRHO, KIAA0283))	624	1E-177
	AA	AC50299.1	protein tyrosine phosphatase sigma	437	1E-120
	AN.		002841.2 protein tyrosine phosphatase, receptor type, sigma, isoform 1 precursor; protein tyrosine	437	1E-120
			phosphatase PTPsigma		
	AA	AC41749.1	protein tyrosine phosphatase delta	436	1 e-120
	N.	569077.1	NP_569077.1 protein tyrosine phosphatase, receptor type, D isoform 4 precursor; protein tyrosine phosphatase,	436	1E-120
			receptor type, delta polypeptide; protein tyrosine phosphatase delta		
	NP		569075.1 protein tyrosine phosphatase, receptor type, D isoform 2 precursor; protein tyrosine phosphatase,	436	1E-120
			receptor type, delta polypeptide; protein tyrosine phosphatase delta		
	<u>N</u>		569076.1 protein tyrosine phosphatase, receptor type, D isoform 3 precursor; protein tyrosine phosphatase,	436	1E-120
			receptor type, delta polypeptide; protein tyrosine phosphatase delta		
	CA	AA38068.1	CAA38068.1 protein-tyrosine phosphatase	436	1E-120
	N.	570924.1	protein tyrosine phosphatase, receptor type, sigma, isoform 2 precursor; protein tyrosine	434	1E-120
·			phosphatase PTPsigma		
			570923.1 protein tyrosine phosphatase, receptor type, sigma, isoform 3 precursor; protein tyrosine	434	1E-120
			phosphatase PTPsigma		
	NP 	570925.1	NP_570925.1 protein tyrosine phosphatase, receptor type, sigma, isoform 4 precursor; protein tyrosine	434	1E-120
			phosphatase PTPsigma		
	AA	AC50567.1	PTPsigma	432	1E-119
	AA	AD09360.1	AAD09360.1 PTPsigma-(brain)	432	1E-119
	AN.	569707.1	protein tyrosine phosphatase, receptor type, F, isoform 2 precursor; protein tyrosine phosphatase,	431	1E-119
			receptor type, F polypeptide; receptor-linked protein-tyrosine phosphatase LAR; Leukocyte		
			antigen-related tyrosine phosphatase; LCA-homolog		

10

	TDHULK	leukocyte antigen-related protein precursor	431	1E-119
	ILARA	Chain A, Crystal Structure Of The Tandem Phosphatase Domains Of Rptp Lar	425	1E-117
	AAC62834.1	PTPsigma [AA 524- 1926]	420	1E-115
	IRPMB	Chain B, Human Receptor Protein Tyrosine Phosphatase Mu, Domain 1	416	1E-114
NM_019935 Mm.38323 U:2.52 (5to11)	11) 014753	Putative transcription factor Ovo-like 1 (hOvo1)	468	1E-130
NP 064319.1				
	NP_004552.1	004552.1 OVO-like 1 binding protein; putative transcription factor OVO-like 1; ovo (Drosophila) homolog-	367	1E-100
		like 1		
	NP_067043.1	zinc finger protein 339; ovo-like 2 (Drosophila)	275	3E-72
	Q9BRP0	Zinc finger protein 339	271	2E-71
	CAB45151.1	hypothetical protein, similar to (AF134804) putative zinc finger transcription factor OVO1	238	3E-61
NM_033174 Mm.195990 U:2.51 (YtoO)		B/B' protein (AA 1-231)	240	3E-62
NP 149409.1				
	AAA60151	snRNP polypeptide B.	240	3E-62
	CAB57868	snRNP B' protein	240	3E-62
NM_008714 Mm.31255 U.2.5 (5to19)	(9) P46531	Neurogenic locus notch homolog protein 1 precursor (Notch 1) (hN1) (Translocation-associated	646	0
NP 032740.1		notch protein TAN-1)		
	A40043	notch protein homolog TAN-1 precursor	4628	0
	AAA60614.1	TAN1	4482	0
	NP_077719.2	077719.2 notch 2 preproprotein	2628	0
	AAG37073.1	G37073.1 NOTCH2 protein	2627	0
	Q04721	Neurogenic locus notch homolog protein 2 precursor (Notch 2) (InN2)	2627	0
	AAC14346.1	Notch3	2065	0
,	NP_000426.1	000426.1 Notch homolog 3	2065	0
	AAC15789.1	Notch 3	2065	0

. 15

	4; Notch (Drosophila) homolog 1023 0	. 1023 0	0 878 0	521 1E-146	521 IE-146	521 IE-146	521 1E-146	520 1E-145	510 1E-142	507 IE-141	506 1E-141	506 1E-141	506 IE-141	506 1E-141	504 1E-141	500 1E-139	0 865 0		-alpha-hydroxylase; cholesterol 861 0		342 9E-93	342 9E-93	298 2E-79	279 7E-74	259 1E-67	246 SE-64	246 9E-64	
697	P_004548.1 Notch homolog 4 (Drosophila); Notch, drosophila, homolog of, 4; Notch (Drosophila) homolog	notch4	NOTCH4	transmembrane protein Jagged 1	Jaggedl	jagged 1 precursor; jagged 1; jagged1 (Alagille syndrome)	transmembrane protein Jagged	fibrillin-2 precursor	NP_001990.1 fibrillin 2	unnamed protein product	fibrillin 1; Fibrillin-1	fibrillin	fibrillin 1 precursor - human (fragment)	fibrillin	similar to fibrillin	BAB20317.1 notch4	cholesterol 7-alpha hydroxylase		cytochrome P450, subfamily VIIA, polypeptide 1; cholesterol 7-alpha-hydroxylase; cholesterol	7 alpha-monooxygenase	oxysterol 7alpha-hydroxylase	oxysterol 7alpha-hydroxylase	sterol 12-alpha-hydroxylase CYP8B1	sterol 12-alpha hydroxylase CYP8B1	CYP7	prostacyclin synthase	prostacyclin synthase	
	NP_004548.1	AAC63097.1	BAA13116.1	AAB39007.1	AAC51731.1	NP_000205.1	AAC51323.1	A54105	NP_001990.1	BAB55419.1	NP_000129.1	1713408A	A47221	CAA45118.1	XP_034890.4	BAB20317.1	AAA58435		NP_000771		AAC95426	AAD20021	AAD19877	AAC63037	AAA61350	BAA11910.	BAA07343	
																	U:2.47 (YtoM)											
													·				NM_007824 Mm.57029	NP 031850.1										

		AAG31785	prostacyclin synthase	245	2E-63
		AAG31784	prostacyclin synthase	245	2E-63
		AAG31783	prostacyclin synthase	242	2E-62
NM_019640 Mm.200516 U:2.47 (YtoM)	U:2.47 (YtoM)	CAB63033	dJ353E16.1 (phosphatidylinositol transfer protein beta)	929	1E-157
NP 062614.1					
		AAH31427	Similar to phosphotidylinositol transfer protein, beta	549	1E-155
		Q00169	Phosphatidylinositol transfer protein alpha isoform (PtdIns transfer protein alpha) (PtdInsTP) (PI-	461	1E-129
			TP-alpha).		
		BAA06276	phosphatidylinositol transfer protein	457	1E-128
			NIR2	229	3E-59
		CAA67224	homologue of Drosphila retinal degeneration B gene	229	3E-59
		AAH22230	Unknown (protein for MGC:21235)	229	3E-59
		BAA95981	KIAA1457 protein	222	6E-57
		968S90 dN	PYK2 N-terminal domain-interacting receptor 3; KIAA1457 protein; likely ortholog of mouse	222	6E-57
·			retinal degeneration B2 homolog (Drosophila) (Rdgb2)		
NM_019992 Mm.38392	U:2.47 (YtoO)	NP_036240	BCR downstream signaling 1	465	1E-130
NP 064376.1					
NM_011076 Mm.16086	U:2.45 (YtoO)	AAA59576	P glycoprotein.	2029	0
		AAAS9575	P-glycoprotein	2026	0
		NP_000918	ATP-binding cassette, sub-family B (MDR/TAP), member 1; P glycoprotein 1/multiple drug	2023	0
			resistance 1; P-glycoprotein-1/multiple drug resistance-1; multidrug resistance 1		
		DVHU3	multidrug resistance protein 3	1702	0
		NP_061337	ATP-binding cassette, subfamily B, member 4 isoform B; P glycoprotein 3/multiple drug	1697	0
-			resistance 3; P-glycoprotein-3/multiple drug resistance-3; multiple drug resistance 3		
		NP_061338	ATP-binding cassette, subfamily B, member 4 isoform C; P glycoprotein 3/multiple drug	1615	0

WO 2005/000335 PCT/US2004/017322

			231		
			resistance 3; P-glycoprotein-3/multiple drug resistance-3; multiple drug resistance 3		
		AAD28285	bile salt export pump	1172	0
	,	NP_003733	ATP-binding cassette, sub-family B (MDR/TAP), member 11; ABC member 16, MDR/TAP	1171	0
			subfamily; progressive familial intrabepatic cholestasis 2; bile salt export pump		ı
		AAC77455	bile salt export pump	1170	0
NM_009345 Mm.25620	U:2.43 (YtoO)	AAH12920.	Similar to deoxynucleotidyltransferase, terminal	771	0
NP 033371.1					
		NP_004079	deoxynucleotidyltransferase, terminal; Terminal deoxynucleotidyltransferase	167	0
		AAA36726	terminal transferase.	765	0
		AAA61136	terminal deoxynucleotidyltransferase	698	1E-100
		NP_037416.	polymerase (DNA directed), mu; polymerase (DNA-directed), mu	333	8E-90
NM_010225 Mm.6260	U:2.42 (5to11)	NP_001443.1	001443.1 forkhead box F2; forkhead (Drosophila)-like 6	521	1E-146
NP 034355.1					
		2208384B	transcription factor FREAC-2	208	1E-142
3		NP_001442.1	001442.1 forkhead box F1; forkhead (Drosophila)-like 5; Forkhead, drosophila, homolog-like 5; forkhead-	251	2E-65
		ŕ	related activator 1		
AK009815 Mm.29201	1 U:2.42 (YtoO)	AAH12098	Similar to glucan (1,4-alpha-), branching enzyme 1 (glycogen branching enzyme, Andersen	1343	0
BAB26519.1			disease, glycogen storage disease type IV)		
		A46075	1,4-alpha-glucan branching enzyme (EC 2.4.1.18)	1341	0
NM_011178 Mm.2364	U:2.42 (YtoM)	P24158	Myeloblastin precursor (Leukocyte proteinase 3) (PR-3) (PR3) (AGP7)(Wegener's autoantigen)	329	3E-89
NP 035308.1			(P29) (C-ANCA antigen) (Neutrophil proteinase 4) (NP-4).		
		CAA39203	proteinase 3	329	3E-89
		AAB59493	proteinase 3	329	3E-89
		PRHU3	proteinase 3 (BC 3.4.21) precursor [validated]	326	2E-88
		CAA39597	proteinase 3 precursor (AA 238) (1 is 2nd base in codon)	326	2E-88
		1FUJC	Chain C, Pr3 (Myeloblastin).	325	88- 3 9

Q)	
2	

			parameter and the second secon			
			AAA59558	myeloblastin.	324	7E-88
			CAA39598	alternative reading frame (AA 215)	294	8E-79
			ELHUL	leukocyte elastase (EC 3.4.21.37) precursor [validated]	256	2E-67
			CAA29300	medullasin N-term.	253	2E-66
			BAA00128	medullasin	251	6E-66
			1H1BB	Chain B, Crystal Structure Of Human Neutrophil Elastase Complexed With An Inhibitor	. 251	6E-66
				(Gw475151).		
			AAB19409	medullasin [human, bone marrow, Peptide, 238 aa].	251	6E-66
			IPPFE	Chain E, Human Leukocyte Elastase (Hle) (Neutrophil Elastase (Hne)) (E.C.3.4.21.37) Complex	251	6E-66
				With The Third Domain Of Turkey Ovomucoid Inhibitor (Omtky3).		
			AAD21524	proteinase 3	251	6E-66
			1B0FA	Chain A, Crystal Structure Of Human Neutrophil Blastase With Mdl 101, 146	250	1E-65
			HINEE	Chain E, Human Neutrophil Elastase (HNE) (E.C.3.4.21.37) (Also Referred To As Human	249	2E-65
				Leucocyte Elastase (HLE)) Complex With Methoxysuccinyl-Ala-Ala-Pro-Ala Chloromethyl		
				Ketone (MSACK)		••••
	·		1FY3A.	Chain A, [g175q]hhp, A Mutant Of Human Heparin Binding Protein (Cap37).	198	8E-50
			1AES	Human Heparin Binding Protein	198	8E-50
			AAB59353	azurocidin	198	8E-50
			1617124A	cationic antimicrobial protein CAP37	198	8E-50
	•		CAA41601	azurocidin	198	.8E-50
NM_007760 Mm.20396		U:2.41 (5to7)	NP_000746.2	000746.2 carnitine acetyltransferase precursor, isoform 1	1151	0
NP 031786.1						
			P43155	Carnitine O-acetyltransferase (Carnitine acetylase) (CAT)	1139	0
			CAA55359.1	carnitine acetyltransferase	1124	0
			NP_003994.2	carnitine acetyltransferase isofоrm 2	1117	0
			NP_659006.1	659006.1 carnitine acetyltransferase precursor, isoform 3	514	1E-144
			P28329	Choline O-acetyltransferase (CHOACTase) (Choline acetylase) (ChAT)	470	1E-131
			AAK08951.1	AAK08951.1 choline acetyltransferase isoform S	470	IE-131

S

0

러

N

(T

Ŋ

				.293	[3	3,50
•			NP_065574.1	065574.1 choline acetyltransferase isoform 2; acetyl CoA:choline O-acetyltransferase	468	1E-130
			AAK08952.1	choline acetyltransferase isoform R	468	1E-130
			NP_066266.1	NP_066266.1 choline acetyltransferase isoform 1; acetyl CoA:choline O-acetyltransferase	465	1E-129
			T01786	choline acetyltransferase - human (fragment).	409	1E-113
NM_008151 Mm.4720	n.4720	U:2.39 (7to19)	NP_005279.1	005279.1 G protein-coupled receptor 12	290	1E-167
NP 032177.1					3,5,7	r.
			NP_005272.1 C	G protein-coupled receptor 3; adenylate cyclase constitutive activator	354	7E-90
			NP_005275.1	NP_005275.1 G protein-coupled receptor 6	348	1E-94
			AAA64594.1	G protein-coupled receptor	303	3E-81
NM_010846 Mm.33996	n.33996	U:2.39 (YtoO)	NP_002453	myxovirus resistance protein 1; interferon inducible protein p78; interferon-regulated resistance	794	0
NP 034976.1				GTP-binding protein		
			AAA36337	interferon-induced Mx protein	791	0
			BAC04017	unnamed protein product	735	0
			AAH14222	Similar to myxovirus (influenza) resistance 1, homolog of murine (interferon-inducible protein	1 710	0
-			,	p78)		
			B33481	interferon-induced viral resistance protein MxB	989	0
			AAA36459	p78-related protein	989	0
			AAC08451	MX2	376	1E-102
			AAC08448	MX2	311	4E-83
			JC4305	dynamin II - human	228	5E-58
			P50570	Dynamin 2	226	2E-57
			NP_004936	dynamin 2; Dynamin II	226	2E-57
			B40671	dynamin, internal form 2, short C-terminal form	225	4E-57
			AAA02803	dynamin	225	4E-57
			A40671	dynamin, internal form 1, long C-terminal form	223	1E-56
			090016	Dynamin 3 (Dynamin, testicular) (T-dynamin).	219	2E-55
			BAA74843	KIAA0820 protein	219	2E-55
			CAB66647	bypothetical protein	217	8E-55
·						

	0 0 2 0 8 Min. 3925	U:2.39 (5to11)	NP_002952.1	302952.1 S100 calcium-binding protein A4; 18A2; 42A; S100 calcium-binding protein A4 (calciun protein,	159	2E-38
BAA00148.1			1	calvasculin, metastasin, murine placental homolog); malignant transformation suppression 1		
NM_012050 Mm.139817 U:2.38 (YtoO)	Vlm.139817	U:2.38 (YtoO)	AAH46356	osteomodulin	<i>L</i> 99	0
NP 036180.1						
			AAH32667	keratocan	245	2E-63
			NP_002716	proline arginine-rich end leucine-rich repeat protein	232	1E-59
			AAH35281	Similar to fibromodulin	209	8E-53
			CAA51418	fibromodulin	208	2E-52
			NP_002014	fibromodulin precursor	202	1E-50
		-	AAA85268	lunican	200	7E-50
7			P51884	Lumican precursor (Keratan sulfate proteoglycan lumican) (KSPG lumican).	200	7E-50
NM_019748 Mm.29698	Mm.29698	U:2.38 (YtoM)	AAH18271	SUMO-1 activating enzyme subunit 1	594	1E-168
NP 062722.1			A A D 1 2 7 8 5	SIMO 1 activoting angume E1 N gribunit	501	112 167
			AAU 12705	סטונס-ז-מטונעמוווון פודעוווים ביו זא פונטמווים	171	11-11
			AAF29104	HSPC140	587	1E-166
- NM_009676 Mm.26787	Mm.26787	U:2.36 (5to7)	BAB40305.1	BAB40305.1 aldeyde oxidase	2204	0
NP 033806.1						
			Q06278	Aldehyde oxidase	2174	0
			NP_001150.2	aldehyde oxidase 1	2171	0
			P47989	Xanthine dehydrogenase/oxidase [Includes: Xanthine dehydrogenase (XD); Xanthine oxidase	1262	0.
				(XO) (Xanthine oxidoreductase)]		
			AAA75287.1	A75287.1 xanthine dehydrogenase	1261	0
			NP_000370,1	000370.1 xanihene dehydrogenase; xanthine oxidase; xanthine dehydrogenase	1255	0 -
•			XP_002472.7	002472.7 similar to Xanthine dehydrogenase/oxidase	915	0
			XP 172060.1	172060 I similar to ALDEHYDE OXIDASE HOMOLOG-1~data source: SPTR. source lev: O9ESH4	838	C

5

7(

 \dashv

 $\bar{\alpha}$

295

			232		
			evidence:ISS~putative		
NM_008273 Mm.4931	U:2.36 (5to11)	P31277	Ноптеорох ргоtеіп Нох-D11 (Нох-4F)	379	1E-104
NP 032299.1					
		NP_067015.2	067015.2 homeo box D11; homeo box 4F; Hox-4.6, mouse, homolog of, homeobox protein Hox-D11	377	1E-103
		NP_005514.1	005514.1 homeobox protein A11; homeobox protein HOXA11; homeo box 11	226	4E-58
NM_009773 Mm.29133	U:2.35 (MtoO)	C19118	MAD3-like protein kinase	1439	0
NP 033903.1					
		AAL10712	budding uninhibited by benzimidazoles 1 beta	1437	0
		AAC33435	mitotic checkpoint protein kinase	1437	0
		AAC06260	mitotic checkpoint kinase Mad3L	1437	0
		AAH18739	budding uninhibited by beuzimidazoles 1 (yeast homolog), beta	1436	0
		AAC23736	protein kinase	1436	0
NM_030127 Mm.41957	U:2.32 (YtoO)	P83110	Probable serine protease HTRA3 precursor	771	0
NP 084403.1				1	
		NP_002766.	protease, scrine, 11	451	1E-125
		BAC11470	unnamed protein product	383	1E-105
		AAC97211	serine protease	371	1E-101
		AAB94569	serine protease	307	6E-82
		1LCYA	Chain A, Crystal Structure Of The Mitochondrial Serine Protease Htra2.	306	1E-81
		AAF66597.	serine protease HtrA2-p7	258	3E-67
NM_011086 Mm.38370 NP 035216.1	0 U:2.32 (5to11)	BAC03674.1	C03674.1 unnamed protein product	1860	0
		Q9Y2I7	FYVE finger-containing phosphoinositide kinase (1-phosphatidylinositol-4-phosphate 5-kinase)) 1070	0
			(PIP5K) (PtdIns(4)P-5-kinase) (p235)		
		XP_028867.1	028867.1 similar to FYVE finger-containing phosphoinositide kinase (1-phosphatidylinositol-4-phosphate 1032	1032	0
			kinase) (P1P5K) (PtdIns(4)P-5-kmase) (p235)		

				-	
:		NP_689884.1	689884.1 hypothetical protein MGC40423	572	1E-161
		XP_114261.1	114261.1 similar to FYVE finger-containing phosphoinositide kinase (1-phosphatidylinositol-4-phosphate	e 472	1E-131
	٠		kinase) (PIP5K) (PtdIns(4)P-5-kinase) (p235)		
V00795	U:2.3 (5to19)	AAB28159.1	AAB28159.1 anti-colorectal carcinoma heavy chain	327	3E-89
CAA24176.1					
		XP_208769.1	208769.1 similar to Ig gamma-2 chain C region	234	4E-61
	-	AAB21082.1	AAB21082.1 Ig gamma 2 H chain BUR	234	4E-61
		AAG00910.2	AAG00910.2 recombinant IgG2 heavy chain	233	5E-61
		CAC12842.1	immunoglobulin heavy chain constant region	233	5E-61
		P01859	Ig gamma-2 chain C region	233	SE-61
		AAN76042.1	immunoglobulin gamma 2 heavy chain constant region	233	5E-61
		AAN76043.1	AAN76043.1 immunoglobulin gamma 2 heavy chain constant	233	5E-61
		BAC04996.1	BAC04996.1 unnamed protein product	232	1E-60
		AAG00909.1	AAG00909.1 recombinant IgG1 heavy chain	232	2E-60
		869339	Ig heavy chain V region precursor -	232	2E-60
		AA017821.1	AAO17821.1 anti-1abies SO57 inununoglobulin heavy chain	232	2E-60
		CAA75032.1	CAA75032.1 immunoglobulin lambda heavy chain	232	2E-60
		AAH16381.1	Similar to immunoglobulin heavy constant gamma 3 (G3m marker)	232	2E-60
		1605217A	Ig gammal	232	2E-60
		BAC04226.1	unnamed protein product	232	2E-60
•		AAH41037.1	Siniilar to immunoglobulin heavy chain 4 (serum IgG1)	232	2E-60
·		BAC05203.1	unnamed protein product	232	2E-60
		BAC05009.1	unnamed protein product	232	2E-60
	-	BAC05186.1	BAC05186.1 lumamed protein product	232	2E•60
		AA017823.1	anti-rabies SOJA immunoglobulin heavy chain	232	2E-60
		CAC10265.1	immunoglobulin heavy chain	231	2E-60
		CAC10256.1	innnunoglobulin heavy chain	231	2E-60
		CAC10259.1	immunoglobulin heavy chain	231	2E-60
		CAC10242.1	immunoglobulin heavy chain	231	3E-60

				1
	CAC20455.1		230	4E-60
	GHHU	lg gamma-1 chain C region - human	229	7E-60
	CAC20454.1	immunoglobulin heavy chain constant region gamma 1	229	7E-60
	AAC82527.1	AAC82527.1 immunoglobulin gamna-1 heavy chain constant region	229	7E-60
	AAL96263.1	immunoglobulin gamma-1 heavy chain constant region	229	7E-60
	AAK58686.2	K58686.2 factor VII active site mutant immunoconjugate	229	7E-60
	AAH19046.1	AAH19046.1 Similar to immunoglobulin heavy constant gamma 3 (G3m marker)	229	7E-60
	AAH19337.1	H19337.1 Similar to immunoglobulin heavy constant gamma 3 (G3m marker)	229	7E-60
	AAA02914.1	1gG	229	7E-60
	BAC05017.1	unnamed protein product	229	7E-60
	BAC04208.1	BAC04208.1 unnamed protein product	229	7E-60
	AAH14258.1	Similar to immunoglobulin heavy constant gamma 3 (G3m marker)	229	7E-60
	AAH25314.1	AAH25314.1 Similar to immunoglobulin heavy constant gamma 3 (G3m marker)	229	7E-60
	AAH18747.1	Similar to immunoglobulin heavy constant gamma 3 (G3m marker)	229	7E-60
	AAH37361.1	AAH37361.1 similar to inmunoglobulin heavy constant gamma 3	229	7E-60
	BAC05021.1	unnamed protein product	229	7E-60
	BAC05016.1	BAC05016.1 unnamed protein product	229	7E-60
	BAC05018.1	unnamed protein product	229	7E-60
	AAH06402.1	AAH06402.1 Similar to immunoglobulin heavy constant gamma 3 (G3m marker)	229	7E-60
	AAH14667.1	AAH14667.1 Similar to innunnoglobulin heavy constant gamna 3 (G3m marker)	229	7E-60
	AAH26038.1	AAH26038.1 Similar to immunoglobulin heavy constant gamma 3 (G3m marker)	229	7E-60
	BAC05022.1	unnamed protein product	229	7E-60
	BAC05014.1	unnamed protein product	229	7E-60
NM_026189 Mm.6825 U:2.29 (5to11)		BAB21797.1 KIAA1706 protein	616	0
NP 080465.1				
	XP_166595.	XP_166595.1 similar to RIKEN cDNA 2310005P05	616	0
	BAB55076.	BAB55076.1 unnamed protein product	919	0

Ц

 Γ

<u>5</u>

20

				298		
NM_008673 Mm.14125		U:2.29 (MtoO)	CAC01128	arylamine N-acetyltransferase-1	462	1E-129
NP 032699.1						
			AAG23842	arylamine N-acetyltransferase 1	461	1E-129
			AAC24712	N-acetyltransferase-1	460	1E-128
			AAB84384	mutant arylamine N-acetyltransferase	457	1E-127
			BAA14095	arylamine N-acetyltransferase	454	1E-126
			BAA01641.	arylamine N-acetyltransferase	434	1E-120
			B34585	arylamine N-acetyltransferase (EC 2.3.1.5) 2	433	1E-120
			AAG34181	N-acetyltransferase	432	1E-120
			AAA98976	arylamine N-acetyltransferase	432	1E-120
			AAA64584	arylannine N-acetyltransferase	432	1E-120
			AAH15878	N-acetyltransferase 2 (arylamine N-acetyltransferase)	432	1E-120
			AAC03773	N-acetyltransferase	431	1E-119
			AAC14117	N-acetyltransferase	431	1E-119
			AAK51710	N-acetyltransferase 2	430	1E-119
			AAK51711	N-acetyltransferase 2	429	1E-119
NM_013739 Mm.33910	133910	U:2.29 (YtoM)	NP_079148	hypothetical protein FLJ22570	603	1E-172
NP 038767.1					╛	
NM_016922 Mm.103414 U.2.29 (YtoO)	103414	U:2.29 (YtoO)	666660	Galactosylceramide sulfotransferase (GalCer sulfotransferase) (Cerebroside sulfotransferase) (3'-	- 678	0
NP_058618.1		-		phosphoadenylylsulfate: galactosylceramide 3'-sulfotransferase) (3'-phosphoadenosine-		
				S'phosphosulfate: GalCer sulfotransferase).		
			AAH14649	Similar to cerebroside(3'-phosphoadenylylsulfate: galactosylceramide 3') sulfotransferase	678	0
	,		NP_071417	glycoprotein beta-Gal 3'-sulfotransferase	233	6E-60
			AAK01945	beta-galactose-3-O-sulfotransferase 3	224	4E-57
			AAK73365	Galbela 1-3 GalNAc 3'-sulfotransferase	211	3E-53
			CAD38686	hypothetical protein	211	3E-53
			BAB13977	unnamed protein product	210	4E-53

Ŋ

20

-			299		
		AAH12976	beta-galactose-3-O-sulfotransferase, 4	209	7E-53
		AAL55759	ипкпочи	207	5E-52
A F 0 4 7 7 2 5 Mm.42100	U:2.28 (5to11)	NP_000763.1	000763.1 cytochrome P450, subfamily IIC (mephenytoin 4-hydroxylase), polypeptide 18; cytochrome P450,	704	0
AAD13720.1	*. -		subfamily IIC (mephenytoin 4-hydroxylase), polypeptide 17; microsomal monooxygenase;		
			flavoprotein-linked monooxygenase		
-		P33260	Cytochrome P450 2C18 (CYPIIC18) (P450-6B/29C)	704	0
		NP_000760.1	NP_000760.1 cytochrome P450, subfamily IIC (mephenytoin 4-hydroxylase), polypeptide 19; mephenytoin 4-	- 683	0
	·		hydroxylase; microsomal monooxygenase; xenobiotic monooxygenase; flavoprotein-linked		
			monooxygenase		
		P10632	Cytochrome P450 2C8 (CYPIIC8) (P450 form 1) (P450 MP-12/MP-20) (P450 IIC2) (S-	- 681	0
	ŕ		mephenytoin 4-hydroxylase)	· , · · -	
		AAH20596	Unknown (protein for MGC:22146)	089	0
		NP_000762.2	000762.2 cytochrome P450, subfamily IIC, polypeptide 9; cytochrome P450, subfamily IIC (mephenytoin	629	0
			4-hydroxylase), polypeptide 10; mephenytoin 4-hydroxylase; microsomal monooxygenase;		
			xenobiotic monooxygenase; flavoprotein-linked monooxygenase		
		AAB23864.2	B23864.2 cytochrome P-450	629	0
		AAA52161.1	cytochrome P-450 S-mephenytoin 4-hydroxylase	629	0
		BAA00123.1	BAA00123.1 cytochrome P-450	629	0
		NP_000761.2	NP_000761.2 cytochrome P450, subfamily IIC, polypeptide 8 isoform 1; mephenytoin 4-hydroxylase,	, 678	0
			microsomal monooxygenase; xenobiotic monooxygenase; flavoprotein-linked monooxygenase;		
			P450 form 1		
		866382	cytochrome P450 2C8 - human	229	0
		AAB35292.1	.B35292.1 cytochrome P450 arachidonic acid epoxygenase isoform, Cyp 2C8 [human, kidney, Peptide	£ 677	0
			Partial, 485 aa]		
		AAA52160.1	AAA52160.1 cytochrome P-450 S-mephenytoin 4-hydroxylase	677	0
		F38462	S-mephenytoin 4'-hydroxylase (EC 1.14.14) cytochrome P450 2C19	929	0
		P11713	Cytochrone P450 2C10 (CYPIIC10) (P450 MP-8) (S-mephenytoin 4-hydroxylase) (P-450MP)) 674	0
		AAA52157.1	AAA52157.1 cytochrome P-450 S-mephenytoin 4-hydroxylase	674	0

5

⊃ --i

			ſ	, 000		
			1506290A	cytochrome P450	674	0
			152418	cytochrome P450	640	0
NM_058212 Mm.45029	5029	U:2.26 (5to 19)	AAH26305.1	AAH26305.1 cerebellun D4	583	1E-165
NP 478119.1						
		-	NP_006259,1	006259.1 requiem; ubi-d4, apoptosis response zinc finger protein	335	1E-90
			NP_036206.1	cer-d4 (mouse) homolog; hypothetical protein FLJ14079	290	4E-77
			NP_004638.1	004638.1 Neuro-d4 (rat) homolog; D4, zinc and double PHD fingers, family 1	277	3E-73
NM_018861 Mm.6379	5379	U:2.25 (YtoM)	AAA19438.1	AAA19438.1 neutral amino acid transporter	9/9	0
NP 061349.1						
			NP_003029.2 solute	solute carrier family 1, member 4; glutamate/neutral amino acid transporter;	9/9	0
				alanine/serine/cysteine/threonine transporter		
			155389	neutral amino acid transporter	673	0
			BAA94861.1	hASCT1	029	0
			A47131	Na+-dependent neutral amino acid transporter SATT	020	e-179
			AAK77026.1	sodium-dependent neutral amino acid transporter type 2 truncated isoform	365	2E-99
			AAD27806.1	AAD27806.1 sodium-dependent neutral amino acid transporter	365	2E-99
			AAD09814.1	neutral amino acid transporter	365	2E-99
			AAH00062.1	solute carrier family 1 (neutral amino acid transporter), member 5	365	2E-99
			AAC50629.1	neutral amino acid transporter B	365	4E-99
			AAD09812.1	RD114/simian type D retrovirus receptor	361	5E-98
			AAH14403.1	Similar to solute carrier family 1, member 7	324	7E-87
			CAA83507.1	GLAST1	271	4E-71
			AAH37310,1	solute carrier family 1 (glial high affinity glutamate transporter), member 3	271	4E-71
,			838353	glutamate transporter protein - human	792	1E-69
			AAC15754.1	AACI 5754.1 EAT4_HUMAN	761	6E-68
		,	AAH33040.1	AH33040.1 solute carrier family 1 (neuronal/epithelial high affinity glutamate transporter, system Xag),	, 246	1E-63
				member 1		
			P43005	Excitatory amino acid transporter 3 (Sodium-dependent glutamate/asparlate transporter 3)) 246	1E-63
				(Excitatory amino-acid carrier1) (Neuronal and epithelial glutamate transporter).		

2

 $\ddot{}$

. 20

			301		
			excitatory AA transporter:ISOTYPE=3	246	1E-63
		2024230A	neuron-specific Glu transporter III	245	3E-63
	i i	AAC27511.3	neuronal and epithelial glutamate transporter	245	4E-63
		AAA53215.1 HEAAC1	HEAACI	244	7E-63
		NP_004162.1	NP_004162.1 solute carrier family 1, member 2; H.sapiens mRNA for glutamate transporter; glutamate/aspartate	244	1E-62
			transporter II; excitatory amino acid transporter 2; glial high affinity glutamate		
		BAA28706.1	glutamate transporter	243	2E-62
		AAA18900.1	glutamate/aspartate transporter II	243	2E-62
		CAC10342.1	dJ68D18.1.2 (solute carrier family 1 (glial high affinity glutamate transporter) member 2)	243	2E-62
		CAC10343.1	dJ68D18.1.1 (solute carrier family 1 (glial high affinity glutamate transporter) member 2)	243	2E-62
		AAH12119.1	solute carrier family 1 (glutamate transporter), member 7	238	4E-61
		CAC12702.1	C12702.1 bA6J24.1 (solute carrier family 1 (neuronal/epithelial high affinity glutamate transporter, system	237	9E-61
			Xag), member 1)		
		AAB53971.1	retinal glutamate transporter EAATS	234	8E-60
NM_011897 Mm.89982	U.2.25 (YtoO)	AAH15745	sprouty (Drosophila) homolog 2	295	1E-160
NP 036027.1					
		XP_036349	similar to sprouty homolog 1 (Drosophila)	263	4E-69
		043609	Sprouty homolog 1 (Spry-1).	201	3E-50
		NP_005831	sprouty homolog 3; antagonist of FGF signaling	200	4E-50
-					-
A F 1 2 6 8 3 4 Mm.10225	U:2.24 (YtoO)	060437	Periplakin (195 kDa cornified envelope precursor) (190 kDa paraneoplastic pemphigus antigen).	2559	0
AAD20642.1					
		NP_002696	periplakin	2559	0
		AAC17738	195 kDa cornified envelope precursor	2553	, 0
		BAA25494	KIAA0568 protein	1952	0
		AAD00186	envoplakin	629	0
		AAA52288	bullous pemphigoid antigen	280	1E-73
		AAB05427	plectin	278	7E-73

Q15149 Plectin 1 (PLXN) (PCN) (Hemidremssonal protein 1) (IDD).			200		
0 0 8 7 6 2 Mm. 56941 U:2.23 (5to19)		Q15149	Plectin I (PLTN) (PCN) (Hemidesmosomal protein 1) (HD1).	275	8E-72
0 0 8 7 6 2 Mm. 5 6 9 4 1 U. 2. 23 (5 to 19) 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1		NP_056363	bullous pemphigoid antigen 1, isoform 1 cA precursor; dystonin; hemidesmosomal plaque protein	1 259	3E-67
0 0 8 7 6 2 Mm. 56941 U:2.23 (5to19)			bullous pemphigoid antigen 1 isoform 1eB precursor; bullous pemphigoid antigen 1; bullous	259	3E-67
0 0 8 7 6 2 Mm. 5 6 9 4 1 U. 2. 23 (5 to 19) 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1			pemphigoid antigen 1 (230/240kD); dystonin; hemidesmosomal plaque protein		
0 0 8 7 6 2 Mm. 56941 U:2.23 (5to19) 1 32788.1		AAL62062	bullous pemphigoid antigen 1 eB	258	1E-66
0 0 8 7 6 2 Mm. 56941 U:2.23 (5to19)		094833	Bullous pemphigoid antigen 1, isoforms 6/9/10 (Trabeculin-beta) (Bullous pemphigoid antigen)	256	3E-66
0 0 8 7 6 2 Mm. 5 6 9 4 1 U. 2. 23 (5 to 19)]			(BPA) (Hemidésmosomal plaque protein) (Dystonia musculorum protein).		
0 0 8 7 6 2 Mm. 56941 U:2.23 (5to19)		AAL62061	bullous pemphigoid antigen 1 eA	256	3E-66
008762 Mm.56941 U:2.23 (5to19) 1		Q03001	Bullous pemphigoid antigen 1 isoforms 1/2/3/4/5/8 (230 kDa bullous pemphigoid antigen) (BPA)	249	4E-64
008762 Mm.56941 U:2.23 (5to19)			(Hemidesmosomal plaque protein) (Dystonia musculorum protein).		
008762 Mm.56941 U:2.23 (5to19) 132788.1					
	008762 Mm.56941	EZ	olfactory receptor, family 2, subfamily C, member 1	203	1E-141
BAC05729.1 seven transmembrane helix receptor	NP 032788.1				
XP_060573.1 similar to olfactory receptor MOR256-12 AAH30717.1 olfactory receptor, family 2, subfamily C, member 3 BAC05875.1 seven transmembrane helix receptor XP_060575.1 similar to seven transmembrane helix receptor XP_165701.1 similar to olfactory receptor 2B2 (Olfactory receptor 2B2 (Olfactory receptor 2B2 (Olfactory receptor 6-1) (OR6-1) (Hs6M1-10 (Q15062 Olfactory receptor 2B2 (Olfactory receptor 6-1) (AAF98753.1 olfactory receptor 2B2 (Olfactory receptor 6-1) (AAF98752.1 olfactory receptor 2B2 (Olfactory receptor 6-1) (OR6-1) (AAF98752.1 olfactory receptor 6-1) (OR6-1) (OR6		BAC05729.1	seven transmembrane helix receptor	384	1E-105
AAH30717.1 Olfactory receptor, family 2, subfamily C, member 3				383	1E-105
BACU5875.1 seven transmembrane helix receptor		AAH30717.1	olfactory receptor, family 2, subfamily C, member 3	378	1E-104
XP_060575.1 similar to seven transmembrane helix receptor NP_149046.1 olfactory receptor, family 2, subfamily B, member 2 XP_165701.1 similar to Olfactory receptor 2B2 (Olfactory receptor 6-1) (OR6-1) (Hs6M1-10 Q15062 Olfactory receptor 2H3 (Olfactory receptor FAT11) AAF98753.1 olfactory receptor AAF98754.1 olfactory receptor AAF98752.1 olfactory receptor CAC20531.1 olfactory receptor CAC20531.1 olfactory receptor XP_094900.1 similar to Olfactory receptor 2H1 (Hs6M1-16) (Olfactory receptor 6-2) (OR6		BAC05875.1	seven transmembrane helix receptor	377	1E-103
NP_149046.1 Olfactory receptor, faruily 2, subfamily B, member 2 XP_165701.1 Similar to Olfactory receptor 2B2 (Olfactory receptor 6-1) (OR6-1) (Hs6M1-10 Q15062 Olfactory receptor 2H3 (Olfactory receptor-like protein FAT11) AAF98753.1 Olfactory receptor AAF98754. Olfactory receptor AAF98752.1 Olfactory receptor AAF98751.1 Olfactory receptor CAC20531.1 Olfactory receptor CAC20531.1 Olfactory receptor CAC20531.1 Olfactory receptor XP_094900.1 Similar to Olfactory receptor 2H1 (Hs6M1-16) (Olfactory receptor 6-2) (OR6		XP_060575.1		377	1E-103
XP_165701.1 similar to Olfactory receptor 2B2 (Olfactory receptor 6-1) (OR6-1) (Hs6M1-10 Q15062 Olfactory receptor 2H3 (Olfactory receptor-like protein FAT11) AAF98753.1 Olfactory receptor AAF98754. olfactory receptor AAF98752.1 olfactory receptor AAF98751.1 olfactory receptor CAC20531.1 olfactory receptor CAC20531.1 olfactory receptor XP_094900.1 similar to Olfactory receptor 2H1 (Hs6M1-16) (Olfactory receptor 6-2) (OR6			olfactory receptor, family 2, subfamily B, member 2	371	1E-102
Q15062 Olfactory receptor 2H3 (Olfactory receptor-like protein FAT11) O95918 Olfactory receptor AAF98753.1 olfactory receptor AAF98754. olfactory receptor AAF98752.1 olfactory receptor AAF98753.1 olfactory receptor CAC20531.1 olfactory receptor XP_094900.1 similar to Olfactory receptor 2H1 (Hs6M1-16) (Olfactory receptor 6-2) (OR6		XP_165701.1	similar to Olfactory receptor 2B2 (Olfactory receptor 6-1) (OR6-1) (Hs6M1-10)	370	1E-101
AAF98753.1 Olfactory receptor 2H2 (Hs6M1-12) AAF98753.1 olfactory receptor AAF98754.2 olfactory receptor AAF98751.1 olfactory receptor AAF98751.1 olfactory receptor CAC20531.1 olfactory receptor XP_094900.1 similar to Olfactory receptor 2H1 (Hs6M1-16) (Olfactory receptor 6-2) (OR6		Q15062	Olfactory receptor 2H3 (Olfactory receptor-like protein FAT11)	369	1E-101
AAF98753.1 olfactory receptor AAF98754. olfactory receptor AAF98752.1 olfactory receptor AAF98751.1 olfactory receptor CAC20531.1 olfactory receptor XP_094900.1 similar to Olfactory receptor 2H1 (Hs6M1-16) (Olfactory receptor 6-2) (OR6		095918	Olfactory receptor 2H2 (Hs6M1-12)	367	1E-100
AAF98754. olfactory receptor AAF98752.1 olfactory receptor AAF98751.1 olfactory receptor CAC20531.1 olfactory receptor XP_094900.1 similar to Olfactory receptor 2H1 (Hs6M1-16) (Olfactory receptor 6-2) (OR6		AAF98753.1	olfactory receptor	366	1E-100
AAF98751 olfactory receptor AAF98751.1 olfactory receptor CAC20531.1 olfactory receptor XP_094900.1 similar to Olfactory receptor 2H1 (Hs6M1-16) (Olfactory receptor 6-2) (OR6		AAF98754.	olfactory receptor	364	1E-100
AAF98751.1 olfactory receptor CAC20531.1 olfactory receptor XP_094900.1 similar to Olfactory receptor 2H1 (Hs6M1-16) (Olfactory receptor 6-2) (OR6		AAF98752.1	olfactory receptor	. 363	1E-99
CAC20531.1 olfactory receptor XP_094900.1 similar to Olfactory receptor 2H1 (Hs6M1-16) (Olfactory receptor 6-2) (OR6		AAF98751.1	olfactory receptor	363	1E-99
XP_094900.1 similar to Olfactory receptor 2H1 (Hs6M1-16) (Olfactory receptor 6-2) (OR6		CAC20531.1		363	2E-99
		XP_094900.1	XP_094900.1 similar to Olfactory receptor 2H1 (Hs6M1-16) (Olfactory receptor 6-2) (OR6-2) (OLFR42A-	196.	9E-99
9004;14/9026;2)			9004.14/9026.2)		

CAC20478.1 olitatory receptor CAC20478.1 olitatory receptor CAC20478.1 olitatory receptor CAC20478.1 olitatory receptor CAC21440.1 olitatory receptor CAC20480.1 olitatory recepto	303			
XP_173276. similar to Olfactory receptor 213 (Olfactory receptor 6-6) (OR6-6) (H56M1) CAC21440. olfactory receptor CAC21440. olfactory receptor XP_175188. similar to olfactory receptor XP_060530. similar to olfactory receptor MOR256-14 BAC061621. seven transmembrane helix receptor XP_060530. seven transmembrane helix receptor BAC0617. seven transmembrane helix receptor XP_06054. similar to olfactory receptor (mint) 2, subfamily C, member 3 P5817.3 Olfactory receptor 2B6 (H56M1-32) (Olfactory receptor 6-31) (OR6-31) XP_161740. similar to olfactory receptor (mint) 2, subfamily W, member 1 CAC2051.1 olfactory receptor (mint) 2, subfamily W, member 1 CAC2051.2 olfactory receptor (mint) 2, subfamily M, member 1 CAC2051.3 olfactory receptor (mint) 2, subfamily 3, member 1 CAC2051.3 olfactory receptor (mint) 2, subfamily 3, member 1 CAC2051.3 olfactory receptor (mint) 2, subfamily 3, member 1 CAC2051.3 olfactory receptor (mint) 3, subfamily 3, member 1 CAC2051.3 ismilar to olfactory receptor (mint) 4, member 1 CAC20481.1 similar to olfactory receptor (mint) 5, subfamily 3, member 1 CAC20481.1 similar to olfactory receptor (mint) 6, subfamily 3, member 1 CAC20481.1 similar to olfactory receptor (mint) 6, subfamily 3, member 1 CAC20481.1 similar to olfactory receptor (mint) 7, member 1 XP_060578.4 similar to olfactory receptor (mint) 6, (novel 7 transmembrane receptor (hodops) 6, 18 seven transmembrane helix receptor (mint) 7, member 1 CAC20481.1 similar to olfactory receptor (mint) 7, member 1 CAC20481.1 similar to olfactory receptor 8 XP_060578.4 similar to olfactory receptor 8 CAC20481.1 similar to olfactory receptor 8 CAC20481.1 similar to olfactory receptor 8 CAC20481.1 similar to olfactory receptor 8 CAC20481.1 similar to olfactory receptor 8 CAC20481.1 similar to olfactory receptor 8 CAC20481.1 similar to olfactory receptor 8 CAC20481.1 similar to olfactory receptor 8 CAC20481.1 similar to olfactory receptor 8 CAC20481.1 similar to			358	4E-98
CAC21440.1 Olfactory receptor CAC21440.1 Olfactory receptor XP_175188.3 similar to olfactory receptor NP_000001.1 olfactory receptor NP_000501.2 seven transmembrane helix receptor BAC0580.4 similar to olfactory receptor MOR256-14 BAC0580.4.1 seven transmembrane helix receptor BAC0580.4.2 similar to olfactory receptor family 2, subfamily C, member 3 TSP 107140.1 similar to olfactory receptor 133 (Olfactory receptor 6-4) (OR6-4) (Hs6041) NP_112165.1 olfactory receptor 7 CAC20485.1 olfactory receptor 6-4) (OR6-4) (Hs6041) NP_112165.1 olfactory receptor 6-4 CAC20485.1 olfactory receptor 7 CAC20485.1 olfactory receptor 7 CAC20485.1 olfactory receptor 7 CAC20485.1 olfactory receptor 7 CAC20485.1 seven transmembrane helix receptor 7 CAC20485.1 seven transmembrane helix receptor 7 CAC20487.1 seven transmembrane helix receptor 7 CAC20487.1 olfactory receptor 8 CAC20487.3 olfactory receptor 8 CAC20487.4 <th>XP_175276.1 similar to Olfactory receptor 233 (Olfactory receptor 6-0</th> <td></td> <td>358</td> <td>6E-98</td>	XP_175276.1 similar to Olfactory receptor 233 (Olfactory receptor 6-0		358	6E-98
CAC21440.1 Oilactory receptor	CAC20477.1 olfactory receptor		358	6E-98
XP_1751881 similar to olfactory receptor	221440.1		358	7E-98
NP_009091.1 Olfactory receptor, family 2, subfamily H, member 3; Olfactory receptor 2			357	1E-97
XP_060580.4 similar to olfactory receptor BAC06162.1 seven transmembrane helix receptor BAC06162.1 seven transmembrane helix receptor BAC0654.6 similar to olfactory receptor AP 12065.7 Olfactory receptor 2B6 (Hs6M1-32) (Olfactory receptor 6-31) (OR6-31) XP 167140.1 similar to Olfactory receptor 2B6 (Hs6M1-32) (Olfactory receptor 6-31) (OR6-31) XP 167140.1 similar to Olfactory receptor CAC20522.1 olfactory receptor CAC20523.1 olfactory receptor CAC20523.1 olfactory receptor CAC20523.1 olfactory receptor CAC20523.1 olfactory receptor BAC05901.1 seven transmembrane helix receptor CAC20523.1 olfactory receptor CAC20523.1 olfactory receptor CAC2053.1 olfactory receptor CAC2063.1 olfactory receptor CAC2063.2<	NP_009091.1 olfactory receptor, family 2, subfamily H, member 3; O	actory receptor 2	357	1E-97
BAC06162.1 seven transmembrane helix receptor BAC05847.1 seven transmembrane helix receptor XP_060574.6 similar to olfactory receptor family 2, subfamily C, member 3 PS8173 Olfactory receptor 2B6 (Hs6M1-32) (Olfactory receptor 6-31) (OR6-4) (Hs6M1 XP_167140.1 similar to Olfactory receptor 2B3 (Olfactory receptor 6-31) (OR6-4) (Hs6M1 NP_112165.1 olfactory receptor CAC20522.1 olfactory receptor CAC20485.1 olfactory receptor CAC20523.3 olfactory receptor CAC20523.1 olfactory receptor XP_175187.1 similar to olfactory receptor XP_175187.1 similar to olfactory receptor XP_175292.1 similar to olfactory receptor CAC20593.1 olfactory receptor CAC20593.1 olfactory receptor XP_106797.1 olfactory receptor XP_106797.1 similar to olfactory receptor MOR256.3 XP_106797.1 similar to olfactory receptor MOR256.3 XP_2069937.1 similar to olfactory receptor CAA10602.1 olfactory receptor CAA10602.1 olfactory receptor CA	XP_060580.4 similar to olfactory receptor MOR256-14		356	2E-97
BAC05847.1 seven transmembrane helix receptor BAC05847.1 Seven transmembrane helix receptor Exp D60574.6 Similar to olfactory receptor, family 2, subfamily C, member 3 P58173 Olfactory receptor 2B6 (Hs6M1-32) (Olfactory receptor 6-31) (ON6-31)	BAC06162.1 seven transmembrane helix receptor		355	4E-97
XP_060574.6 similar to olfactory receptor, family 2, subfamily C, member 3 P58173 Olfactory receptor 2B6 (Hs6MI-32) (Olfactory receptor 6-4) (OR6-4) (Hs6MI XP_167140.1 similar to Olfactory receptor 2B3 (Olfactory receptor 6-4) (OR6-4) (Hs6MI NP_112165.1 olfactory receptor CAC20522.1 olfactory receptor CAC20533.1 olfactory receptor CAC20533.3 olfactory receptor XP_175187.1 similar to olfactory receptor XP_175187.1 similar to olfactory receptor XP_175292.1 similar to olfactory receptor CAC20497.1 olfactory receptor CAC20503.1 olfactory receptor CAC204097.1 ismilar to olfactory receptor MOR256-12 CAC204097.1 olfactory receptor MOR256-12 XP_17187.1 similar to olfactory receptor MOR256-12 XP_167135.1 similar to olfactory receptor MOR256-13 XP_167135.1 similar to olfactory receptor 8 CAA10602.1 olfactory receptor 8 CAA10602.1 olfactory receptor 8 CAA10602.1 olfactory receptor 8 CAA10602.1 olfactory receptor 8 CAA10602.1 olfactory receptor 8 C	BAC05847.1 seven transmembrane helix receptor		352	5E-96
P58173 Olfactory receptor 2B6 (Hs6M1-32) (Olfactory receptor 6-31) (OR6-31) XP_167140.1 similar to Olfactory receptor 2B3 (Olfactory receptor 6-4) (OR6-4) (Hs6M1 NP_112165.1 olfactory receptor 7B3 (Olfactory receptor 6-4) (OR6-4) (Hs6M1 CAC20522.1 olfactory receptor 7B (AC20682.1 CAC20682.1 olfactory receptor 7B (AC20682.1 XP_175187.1 similar to olfactory receptor 7B (AC20692.1 RAC05901.1 seven transmembrane helix receptor 7B (AC20692.1 AP_172292.1 similar to olfactory receptor 7B (AC20692.1 CAC20497.1 olfactory receptor 7B (AC20692.1 NP_112167.1 olfactory receptor 7B (AC20692.1 NP_112167.1 olfactory receptor 7B (AC20692.1 NP_112167.1 similar to olfactory receptor 7B (AC20692.1 NP_112167.1 olfactory receptor 7B (AC20692.1 NP_112167.	XP_060574.6 similar to olfactory receptor, family 2, subfamily C, me.	lber 3	350	2E-95
XP_167140.1 similar to Olfactory receptor 2B3 (Olfactory receptor 6-4) (OR6-4) (H86M1 NP_112165.1 olfactory receptor CAC20522.1 olfactory receptor CAC20485.1 olfactory receptor CAC20485.1 olfactory receptor CAC20485.1 olfactory receptor CAC20485.1 olfactory receptor XP_175187.1 similar to olfactory receptor XP_175292.1 similar to olfactory receptor CAC20497.1 olfactory receptor CAC20497.1 olfactory receptor CAC20497.1 olfactory receptor CAC20497.1 olfactory receptor XP_04997.1 similar to olfactory receptor XP_04997.1 similar to olfactory receptor XP_04997.1 similar to olfactory receptor XP_04997.1 similar to olfactory receptor XP_04997.1 similar to olfactory receptor XP_04997.1 similar to olfactory receptor XP_04997.1 similar to olfactory receptor XP_04997.1 similar to olfactory receptor XP_04999.1 seven transmembrane CAA10602.1 olfactory receptor XP_04999.2 seven transmembrane XP_04999.3 seven transmembrane XP_049999.3 seven transmembrane XP_049999.3 seven transmembrane XP_049999.3 seven transmembrane XP_049999.3 seven transmembrane XP_0499999.3 XP_04999999999999999999999999999999999999	P58173 Olfactory receptor 2B6 (Hs6M1-32) (Olfactory recepto	6-31) (OR6-31)	350	2E-95
NP_112165.1 olfactory receptor, family 2, subfamily W, member 1	XP_167140.1 similar to Olfactory receptor 2B3 (Olfactory receptor 6	I) (OR6-4) (Hs6M1-1)	348	4E-95
CAC20522.1 Olfactory receptor CAC20485.1 Olfactory receptor CAC20523.1 Olfactory receptor XP_175187.1 similar to olfactory receptor XP_175292.1 similar to olfactory receptor XP_172292.1 similar to olfactory receptor CAC20503.1 olfactory receptor CAC20497.1 olfactory receptor NP_112167.1 olfactory receptor XP_094937.1 similar to olfactory receptor 89 XP_094937.1 similar to olfactory receptor MOR256.3 XP_060578.4 similar to olfactory receptor MOR256.3 XP_060578.4 similar to olfactory receptor BAC0509.1 seven transmembrane helix receptor XP_1670406.2 similar to 573K1.15 (mn17M1-6 (novel 7 transmembrane receptor (rhodops)	NP_112165.1 olfactory receptor, family 2, subfamily W, member 1		347	1E-94
CAC20485.1 olfactory receptor CAC20523.1 olfactory receptor XP_175187.1 similar to olfactory receptor XP_172292.1 similar to olfactory receptor CAC20503.1 olfactory receptor CAC20497.1 olfactory receptor CAC20497.1 olfactory receptor NP_112167.1 olfactory receptor XP_094937.1 similar to olfactory receptor MOR256.3 XP_06937.1 similar to olfactory receptor MOR256.3 XP_167135.1 similar to olfactory receptor MOR256.1 XP_167135.1 similar to olfactory receptor MOR256.1 XP_167136.1 similar to olfactory receptor MOR256.1 CAA10602.1 similar to olfactory receptor RAC05909.1 seven transmembrane helix receptor XP_167046.2 similar to 573K1.15 (mon17M1-6 (novel 7 transmembrane receptor (thodopsi	CAC20522.1 olfactory receptor		347	2E-94
CAC20523.1 olfactory receptor XP_175187.1 similar to olfactory receptor XP_172292.1 similar to olfactory receptor CAC20503.1 olfactory receptor CAC20497.1 olfactory receptor CAC20497.1 olfactory receptor NP_112167.1 olfactory receptor XP_094937.1 similar to olfactory receptor 89 XP_160578.4 similar to olfactory receptor MOR256-3 XP_060578.4 similar to olfactory receptor MOR256-12 CAA10602.1 olfactory receptor 89 BAC05909.1 seven transmembrane helix receptor XP_167046.2 similar to 573K1.15 (mml 7M1-6 (novel 7 transmembrane receptor (thodopsis)	CAC20485.1 olfactory receptor		347	2E-94
XP_175187.1 similar to olfactory receptor BAC05901.1 seven transmembrane helix receptor XP_172292.1 similar to olfactory receptor CAC20503.1 olfactory receptor CAC20497.1 olfactory receptor NP_112167.1 olfactory receptor XP_094937.1 similar to olfactory receptor 89 XP_167135.1 similar to olfactory receptor MOR256-3 XP_060578.4 similar to olfactory receptor MOR256-12 CAA10602.1 olfactory receptor 89 BAC05909.1 seven transmembrane helix receptor XP_167046.2 similar to 573K1.15 (mm17M1-6 (novel 7 transmembrane receptor (rhodopsis))	CAC20523.1 olfactory receptor		346	3E-94
BAC05901.1 seven transmembrane helix receptor XP_172292.1 similar to olfactory receptor CAC20503.1 olfactory receptor CAC20497.1 olfactory receptor NP_112167.1 olfactory receptor XP_094937.1 similar to olfactory receptor MOR256-3 XP_167135.1 similar to olfactory receptor MOR256-12 CAA10602.1 olfactory receptor R9 BAC05909.1 seven transmembrane helix receptor XP_167046.2 similar to 573K1.15 (munl 7M1-6 (novel 7 transmembrane receptor (rhodopsis))	XP_175187.1 similar to olfactory receptor		346	3E-94
XP_172292.1 similar to olfactory receptor CAC20503.1 olfactory receptor CAC20497.1 olfactory receptor NP_112167.1 olfactory receptor XP_094937.1 similar to olfactory receptor MOR256-3 XP_167135.1 similar to olfactory receptor MOR256-3 XP_060578.4 similar to olfactory receptor MOR256-12 CAA10602.1 olfactory receptor 89 BAC05909.1 seven transmembrane helix receptor XP_167046.2 similar to 573K1.15 (mml 7M1-6 (novel 7 transmembrane receptor (rhodopsi	BAC05901.1 seven transmembrane helix receptor		341	9E-93
CAC20503.1 olfactory receptor CAC20497.1 olfactory receptor NP_112167.1 olfactory receptor, family 2, subfamily J, member XP_094937.1 similar to olfactory receptor 89 XP_167135.1 similar to olfactory receptor MOR256-3 XP_060578.4 similar to olfactory receptor MOR256-12 CAA10602.1 olfactory receptor 89 BAC05909.1 seven transmembrane helix receptor XP_167046.2 similar to 573K1.15 (mm17M1-6 (novel 7 transmembrane receptor (rhodopsi	XP_172292.1 similar to olfactory receptor MOR256-12	•	341	9E-93
CAC20497.1 olfactory receptor NP_112167.1 olfactory receptor, family 2, subfamily 3, member XP_094937.1 similar to olfactory receptor 89 XP_167135.1 similar to olfactory receptor MOR256-3 XP_060578.4 similar to olfactory receptor MOR256-12 CAA10602.1 olfactory receptor 89 BAC05909.1 seven transmembrane helix receptor XP_167046.2 similar to 573K1.15 (mml 7M1-6 (novel 7 transmembrane receptor (rhodopsi	CAC20503.1 olfactory receptor		339	3E-92
NP_112167.1 olfactory receptor, family 2, subfamily J, member XP_094937.1 similar to olfactory receptor 89 XP_167135.1 similar to olfactory receptor MOR256-3 XP_060578.4 similar to olfactory receptor MOR256-12 CAA10602.1 olfactory receptor 89 BAC05909.1 seven transmembrane helix receptor XP_167046.2 similar to 573K1.15 (mnn1 ⁷ M1-6 (novel 7 transmembrane receptor (rhodopsi	CAC20497.1 olfactory receptor		337	1E-91
XP_094937.1 similar to olfactory receptor MOR256-3 XP_167135.1 similar to olfactory receptor MOR256-12 CAA10602.1 olfactory receptor 89 BAC05909.1 seven transmembrane helix receptor XP_167046.2 similar to 573K1.15 (mnl 7M1-6 (novel 7 transmembrane receptor (rhodopsis))	NP_112167.1 olfactory receptor, family 2, subfamily J, member		337	1E-91
XP_167135.1 similar to olfactory receptor MOR256-3 XP_060578.4 similar to olfactory receptor MOR256-12 CAA10602.1 olfactory receptor 89 BAC05909.1 seven transmembrane helix receptor XP_167046.2 similar to 573K1.15 (mm17M1-6 (novel 7 transmembrane receptor (rhodopsi))	XP_094937.1 similar to olfactory receptor 89		323	3E-87
XP_060578.4 similar to olfactory receptor MOR256-12 CAA10602.1 olfactory receptor 89 BAC05909.1 seven transmembrane helix receptor XP_167046.2 similar to 573K1.15 (mnl 7M1-6 (novel 7 transmembrane receptor (rhodopsi	XP_167135.1 similar to olfactory receptor MOR256-3		319	4E-86
CAA10602.1 olfactory receptor 89 BAC05909.1 seven transmembrane helix receptor XP_167046.2 similar to 573K1.15 (mm17M1-6 (novel 7 transmembrane receptor (rhodopsi	XP_060578.4 similar to olfactory receptor MOR256-12		305	4E-82
BAC05909.1 seven transmembrane helix receptor XP_167046.2 similar to 573K1.15 (mnn17M1-6 (novel 7 transmembrane receptor (rhodopsi	CAA10602.1 olfactory receptor 89		304	1E-81
XP_167046.2 similar to 573K1.15 (mm17M1-6 (novel 7 transmembrane receptor (rhodopsi	BAC05909.1 seven transmembrane helix receptor		295	6E-79
	XP_167046.2 similar to 573K1.15 (mm17M1-6 (novel 7 transmembra	e receptor (rhodopsin family) (olfactory	295	8E-79
receptor LIKE) protein))	receptor LIKE) protein))			

				304		
			NP_110503.1	110503.1 olfactory receptor, family 5, subfamily V, member 1	292	5E-78
U67189	Mni.181709	Mnr. 181709 U.2.23 (YtoM)	AAM12651	regulator of G protein signalling 16	323	2E-87
AAB50619.1						
			AAC16912	A28-RGS14p	320	2E-86
NM_008008 Mm.57177	Mm.57177	U:2.22 (5to19)	NP_002000.1	002000.1 fibroblast growth factor 7 precursor; keratinocyte growth factor	352	2E-96
NP 032034.1						
NM_013746 Mm.26633	Mm.26633	U:2.22 (YtoO)	NP_067023	pleckstrin homology domain containing, family B (evectins) member 1; PH domain containing	3 450	1E-125
NP 038774.1				protein in retina 1; PH domain containing, retinal 1		
,			AAF21786	KPL1	415	1E-114
			AAF18572	PHR1 isoform 2	364	2E-99
			AAH08075	PH domain containing protein in retina 1	330	7E-89
NM_009613	009613 Mm.89854	U:2.22 (MtoO)	BAA32352.1	BAA32352.1 MDC/ADAM11	1454	0
NP 033743.1						
			075078	ADAM 11 precursor (A disintegrin and metalloproteinase domain 11) (Metalloproteinase-like,	, 1451	0
			(disintegrin-like, and cysteine-richprotein)		
			196591	disintegrin-like metalloproteinase (EC 3.4.24), splice form 2	1345	0
			BAA06670.1	metalloprotease/disintegrin-like protein	1340	0
		1	S38539	disintegrin-like metalloproteinase (EC 3.4.24), splice form 1	1011	0
			BAA06671.1	BAA06671.1 metalloprotease/disintegrin-like protein	1008	0
			AAF22476.2 MDC2	MDC2	825	0
			NP_057435.2	057435.2 a disintegrin and metalloproteinase domain 22 isoform 3 proprotein; MDC2 delta	825	0
			NP_068368.2	068368.2 a disintegrin and metalloproteinase domain 22 isoform 2 proprotein; MDC2 delta	825	0
			AAF73288.1	metalloprotease-like, disintegrin-like, cysteine-rich protein 2 delta	825	0
			NP_068367.1	a disintegrin and metalloproteinase domain 22 isoform 5 proprotein; MDC2 delta	825	0
			NP_004185.1	004185.1 a disintegrin and metalloproteinase domain 22 isoform 4 proprotein; MDC2 delta	821	0
			Q9P0K1	ADAM 22 precursor (A disintegrin and metalloproteinase domain 22)(Metalloproteinase-like,	e, 821	0
						1

ហ

0

15

20

C)

WO 2005/000335 PCT/US2004/017322

	305 disintegrin-like, and cysteine-rich protein 2) (Metalloproteinase-disintegrin ADAM22-3).		
AAD55251.1	metalloproteinase-disintegrin ADAM22-3	821	0
NP_003803.1	a disintegrin and metalloproteinase domain 23 preproprotein	726	0
NP_075525.2	a disintegrin and metalloproteinase domain 19 isoform 1 preproprotein; meltrin beta	359	1E-97
NP_150377.1	a disintegrin and metalloproteinase domain 19 isoform 2 preproprotein; meltrin beta	358	3E-97
CAC20585.1	meltrin-beta/ADAM 19 homologue	358	3E-97
AAG50282.1	metalloprotease-disintegrin meltrin beta	357	7E-97
NP_067673.1	NP_067673.1 a disintegrin and metalloprotease domain 12 isoform 2 preproprotein; A disintegrin and	348	4E-94
	metalloproteinase domain 12(Meltrin-alpha, mouse, homolog of); meltrin alpha		
AAC08703.2	meltrin-S	348	4E-94
AAC08702.2	AAC08702.2 meltrin-L precursor	348	4E-94
NP_079496.1	a disintegrin and metalloproteinase domain 33 isoform alpha preproprotein; disintegrin and	344	4E-93
	reprolysin metalloproteinase familyprotein; metalloprotease disintegrin		
AAM80482.1	a disintegrin and metalloprotease domain 33	344	4E-93
AAF22162.1	disintegrin and metalloproteinase domain 19	338	3E-91
CAC16509.2	dJ964F7.1 (novel disintegrin and reprolysin metalloproteinase family protein)	334	6E-90
NP_694882.1	NP_694882.1 a disintegrin and metalloproteinase domain 33 isoform beta preproprotein; disintegrin and	326	1E-87
	reprolysin metalloproteinase familyprotein; metalloprotease disintegrin		
AAC50403.1	metalloprotease/disintegrin/cysteine-rich protein precursor	320	1E-85
AAH14566.1	a disintegrin and metalloproteinase domain 15 (metargidin)	307	6E-82
AAC50404.1	metargidin preçursor	307	6E-82
AAC51112.1	MDC15 ,~	307	6E-82
AAM49575.1	disintegrin/metalloproteinase domain 9 short protein precursor	302	2E-80
NP_001100.1	a disintegrin and metalloproteinase domain 8 precursor	293	2E-77
NP_055080.1	a disintegrin and metalloproteinase domain 28 isoform 1 preproprotein	292	2E-77
AAD25099.1	metalloprotease disintegrin cysteine-rich protein, transmembrane form MDC-Lm	291	3E-77
BAA03499.2	BAA03499.2 KIAA0021 protein	288	5E-76
NP_003804.1	a disintegrin and metalloproteinase domain 21 preproprotein	270	8E-71

			200		ſ
AF366393 Mm.141563 U.2.21 (YtoM) BAA96065 KIAA1541 protein	3 U:2.21 (YtoM)	BAA96065	KIAA1541 protein	814	5
AAK53703.1					
		AAH31790	protein phosphatase 2 (formerly 2A), regulatory subunit B (PR 52), beta isoform	759	0
		XP 029744	similar to phosphoprotein phosphatase (EC 3.1.3.16) 2A BR gamma regulatory chain - human	734	0
		AAH32954	Similar to protein phosphatase 2 (formerly 2A), regulatory subunit B (PR 52), gamma isoform	731	0
		AAD20987	protein phosphatase 2A BR gamma subunit	729	0
		AAG39636	protein phosphatase 2A1 B gamma subunit IMYPNO1	728	0
	,				
NM_016894 Mm.3272	U:2.2 (5to11)	NP_005846.1	005846.1 receptor (calcitonin) activity modifying protein 1 precursor; calcitonin receptor-like receptor	231	9E-60
NP 058590.1			activity modifying protein 1		
NM_013560 Mm.13849	9 U:2.2 (5to7)	NP_001531.1	001531.1 heat shock 27kDa protein 1; heat shock 27kD protein 1	357	SE-98
NP 038588.1					
		AAH12292.1	AAH12292.1 Similar to heat shock 27kD protein 1	351	5E-96
		AAA62175.1	462175.1 heat shock protein 27	342	2E-93
		AAH14920.1	H14920.1 Unknown (protein for IMAGE;3906970)	333	1E-90
		XP_066514.1	066514.1 similar to Heat shock 27 kDa protein (HSP 27) (Stress-responsive protein 27) (SRP27) (Estrogen-	. 267	7E-71
			regulated 24 kDa protein) (28 kDa heat shock protein		
NM_009127 Mm.140785 U:2.2 (YtoM)	85 U.2.2 (YtoM)	NP_005054	stearoyl-CoA desaturase (delta-9-desaturase)	597	1E-170
NP 033153.1					
		191000	Acyl-CoA desalurase (Stearoyl-CoA desalurase) (Fatty acid desalurase) (Delta(9)-desalurase).	596	1
		AAH05807.	Unknown (protein for MGC:10264)	592	1E-169
		CAA73998	stearoyl CoA desaturase	589	1E-168
		AAF71040	PRO0998	579	1E-165
		154779	stearoyl-CoA desaturase - human (fragment).	377	1E-104
		XP_208174	similar to stearoyl-CoA desaturase (delta-9-desaturase)	273	3E-73
		CAD38567	hypothetical protein	216	6E-56
				_	
NM 019977 Mm.158200 U:2.18 (YtoO)	200 U:2.18 (YtoO)	AAF25204	unknown	540	1E-152
1					

ហ

Ŋ

			307		
NP 064361.1					
		NP_060054	aldehyde reductase (aldose reductase) like 6; similar to mouse aldehyde reductase 6 (renal); myo-	535	1E-151
			inositol oxygenase; kidney-specific protein 32		
		AAK00766	kidney-specific protein 32	528	1E-149
NM_031194 Mm.20839	U:2.18 (MtoO)	AAH22387.1	AAH22387.1 Unknown (protein for MGC:24086)	793	0
NP 112471.1					
		BAB47393.1	organic anion transporter 3	792	٥
		AAD19357.1	AAD19357.1 organic anion transporter 3	675	0
		NP_695008.1	695008.1 solute carrier family 22 member 6 isoform b; renal organic anion transporter 1; para-	474	1E-132
			aminohippurate transporter		
		AAC70004.1	putative renal organic anion transporter 1	474	1E-132
		AAD10052.1	para-aminohippurate transporter	472	1E-132
		NP_004781.2	NP_004781.2 solute carrier family 22 member 6 isoform a; renal organic anion transporter 1; para-	- 472	IE-131
			aminohippurate transporter		
		NP_695011.1	695011.1 solute carrier family 22 member 6 isoform e; renal organic anion transporter 1; para-	- 446	1E-124
			aminohippurate transporter		
J		CAB94830.1	putative organic anion transport	412	1E-114
		CAB97249.1	putative organic anion transport	410	1E-113
		AAK68156.1	RST	356	9E-97
		AAK68155.1	OAT4	354	3E-96
		NP_006663.2	solute carrier family 22 member 7 isoform a; organic anion transporter 2; liver-specific transporter	л 315	1E-84
		BAB68364.1	organic anion transpoter 4 like protein	314	3E-84
		AAG43523.1	organic anion transporter 2	313	5E-84
		AAL12496.1	organic anion transporter 2	310	4E-83
		CAC82910.1	putative integral membrane transport protein	286	9E-76
		BAB85030.1	unnamed protein product	285	2E-75
		BAB83517.1	hUST3	283	6E-75
		BAC11483.1	BAC11483.1 unnamed protein product	262	2E-68

					100
		BAA76350.1	BAA76350.1 organic-cation transporter like 3	770	1E-00
		NP_700357.1	700357.1 urate anion exchanger 1 isoform b; organic anion transporter 4-like; urate transporter 1; solute	226	1E-57
			carrier family 22 member 12		
		BAA36712.1	OCTN2	223	1E-56
		CAA66978.1	organic cation transporter	220	6E-56
		AAH12325.1	AAH12325.1 Similar to solute carrier family 22 (organic cation transporter), member 5	220	8E-56
		NP_003050.2	solute carrier family 22 member 4; organic cation transporter 4; integral membrane transport	218	3E-55
			protein		
		CAA04751.1	extraneuronal monoamine transporter	217	5E-55
		23356.1	OCTN1	216	8E-55
		NP_003048.1	003048.1 solute carrier family 22 member 1 isoform a; organic cation transporter 1	216	1E-54
		CAB95971.1	oct] cds	215	2E-54
		CAA66977.1	CAA66977.1 organic cation transporter	214	4E-54
		CAC39443.1	CAC39443.1 organic cation transporter 3	213	7E-54
		CAC08550.1	CAC08550.1 bA288H12.2 (organic cation transporter, liver)	209	1E-52
		AAK58593.1	organic cation transporter OKB1	207	4E-52
		AAH35973.1	H35973.1 Similar to organic cationic transporter-like 3	207	5E-52
NM_020051 Mm.29275 U	U:2.17 (YtoO)	BAB83913.1	BAB83913.1 putative bHLII transcription factor	250	1E-65
NP 064435.1					
·		Q9NQ33	Achaete-scute homolog 3 (bHLH transcriptional regulator Sgn-1)	248	5E-65
		NP_065697.1	ASCL3	248	5E-65
AF316872 Mm.18539 U	U:2.16 (YtoM)	NP_115785	PTEN induced putative kinase 1; protein kinase BRPK	801	0
AAK28061.1					7
		AAH28215	PTEN induced putative kinase 1	798	0
		AAH09534	Unknown (protein for IMAGE:3891886)	484	1E-135
		BAC11484	unnamed protein product	408	1E-112
NM 054048 Mm.22980 L	U.2.16 (YtoO)	XP 208584	similar to RE1-silencing transcription factor (REST) co-repressor; co-repressor of Rest; Rest co-	129	0

<u>(1)</u>

٠				308		Ĺ
NP 473389.1				repressor		
			NP 775858	hypothetical protein LOC283248	729	0
			A92581	KIAA1343 protein	437	1E-121
			BAA91872	unnamed protein product	437	1E-121
			BAA06686.	KIAA0071	399	1E-110
			NP 055971	REST corepressor; KLAA0071 protein	399	1E-110
			AAH31608	Similar to hypothetical protein MGC28186	356	7E-97
			AAH10608	Unknown (protein for IMAGE:4157757)	330	4E-89
NM_016968	016968 Mm.39300	U:2.16 (YtoO)	AAH26989	Similar to Olg-1 bHLH protein	310	2E-83
NP 058664.1		٠				
			Q8TAK6	Oligodendrocyte transcription factor 1 (Oligo1).	310	2E-83
M62766	Mm.2226	U:2.16 (YtoM)	1HWLD	Chain D, Complex Of The Catalytic Portion Of Human Hmg-Coa Reductase With Rosuvastatin	432	1E-119
AAA37819.1				(Formally Known As Zd4522).		
			IHWLC	Chain C, Complex Of The Catalytic Portion Of Human Hing-Coa Reductase With Rosuvastatin	432	1E-119
				(Formally Known As Zd4522).		
			1HWLB	Chain B, Complex Of The Catalytic Portion Of Human Hrng-Coa Reductase With Rosuvastatin	432	1E-119
				(Formally Known As Zd4522).		
			1HWLA	Chain A, Complex Of The Catalytic Portion Of Human Hmg-Coa Reductase With Rosuvastatin	432	1E-119
				(Formally Known As Zd4522).		
			1HWKD	Chain D, Complex Of The Catalytic Portion Of Human Hmg-Coa Reductase With Atorvastatin.	432	1E-119
			1HWKC	Chain C, Complex Of The Catalytic Portion Of Human Hmg-Coa Reductase With Atorvastatin.	432	
			IHWKB	Chain B, Complex Of The Catalytic Portion Of Human Hing-Coa Reductase With Atorvastatin.	432	1E-119
			1HWKA	Chain A, Complex Of The Catalytic Portion Of Human Hmg-Coa Reductase With Atorvastatin.	432	1E-119
			1HWJD	Chain D, Complex Of The Catalytic Portion Of Human Hmg-Coa Reductase With Cerivastatin	n 432	1E-119
			1HWJC	Chain C, Complex Of The Catalytic Portion Of Human Hmg-Coa Reductase With Cerivastatin	1 432	1E-119
			1HWJB	Chain B, Complex Of The Catalytic Portion Of Human Hmg-Coa Reductase With Cerivastatin	1 432	1E-119
			1HWJA	Chain A, Complex Of The Catalytic Portion Of Human Hmg-Coa Reductase With Cerivastatin	i 432	1E-119

			310		
	MH1	10	Chain D, Complex Of The Catalytic Portion Of Human Hmg-Coa Reductase With Fluvastatin	432	1E-119
	HI	1HWIC (Chain C, Complex Of The Catalytic Portion Of Human Hmg-Coa Reductase With Fluvastatin	432	1E-119
	1H	1HWIB		432	IE-119
	11	1HWIA		432	1E-119
	HI			432	1E-119
	H1		Chain C, Complex Of The Catalytic Portion Of Human Hmg-Coa Reductase With Sinvastatin	432	1E-119
	HI		Chain B, Complex Of The Catalytic Portion Of Human Hmg-Coa Reductase With Simvastatin	432	1E-119
	H1		Chain A, Complex Of The Catalytic Portion Of Human Hmg-Coa Reductase With Simvastatin	432	1E-119
	HI		Chain D, Complex Of The Catalytic Portion Of Human Hmg-Coa Reductase With Compactin	432	1E-119
			(Also Known As Mevastatin).		
	11	1HW8C	Chain D, Complex Of The Catalytic Portion Of Human Hmg-Coa Reductase With Compactin	432	1E-119
			(Also Known As Mevastatin).		
	(IHI)	IW8B	Chain D, Complex Of The Catalytic Portion Of Human Hing-Coa Reductase With Compactin	432	1E-119
			(Also Known As Mevastatin).		
	VHI IHV	LW8A	Chain D, Complex Of The Catalytic Portion Of Human Hrng-Coa Reductase With Compactin	432	1E-119
		٠	(Also Known As Mevastatin).		
	11	1DQAD	Chain D, Complex Of The Catalytic Portion Of Human Hmg-Coa Reductase With Hmg, Coa, And	432	1E-119
			Nadp+.		
•)ΩΙ	OQAC	Chain C, Complex Of The Catalytic Portion Of Human Hmg-Coa Reductase With Hmg, Coa, And	432	1E-119
			Nadp+.		
,	11	1DQAB	Chain B, Complex Of The Catalytic Portion Of Human Hing-Coa Reductase With Hing, Coa, And	432	1E-119
			Nadp+.		
	11	IDQAA	Chain A, Complex Of The Catalytic Portion Of Human Hmg-Coa Reductase With Hmg, Coa, And	432	1E-119
			Nadp+.		
	11	1DQ9D	Chain D, Complex Of Catalytic Portion Of Human Hmg-Coa Reductase With Hmg-Coa.	432	1E-119
	11	1DQ9C	Chain C, Complex Of Catalytic Portion Of Human Hmg-Coa Reductase With Hmg-Coa.	432	1E-119
	11	1DQ9B	Chain B, Complex Of Catalytic Portion Of Human Hmg-Coa Reductase With Hmg-Coa.	432	1E-119
		1DQ9A	Chain A, Complex Of Catalytic Portion Of Human Hmg-Coa Reductase With Hmg-Coa.	432	1E-119
	ΔI	2080	Chain D, Complex Of The Catalytic Portion Of Human Hmg-Coa Reductase With Hing And Coa	432	1E-119

WO 2005/000335 PCT/US2004/017322

		311		
	1DQ8C	Chain C, Complex Of The Catalytic Portion Of Human Hmg-Coa Reductase With Hing And Coa	432	IE-119
	1DQ8B	Chain B, Conpplex Of The Catalytic Portion Of Human Hrng-Coa Reductase With Hrng And Coa	432	1E-119
	1DQ8A	Chain A, Complex Of The Catalytic Portion Of Human Hrng-Coa Reductase With Hrng And Coa	432	1E-119
	NP_000850	3-hydroxy-3-methylglutaryl-Coenzyme A reductase	432	1E-119
	AAG21343	3-hydroxy-3-methylglutaryl-coenzyme A reductase	432	1E-119
	RDHUE	hydroxymethylglutaryl-CoA reductase (NADPH2) (EC 1.1.1.34)	432	1E-119
	AAA52679	3-lıydroxy-3-methylglutaryl coenzyme A reductase	432	1E-119
	P04035	3-hydroxy-3-methylglutaryl-coenzyme A reductase (HMG-CoA reductase).	432	1E-119
	AAH33692	Similar to 3-hydroxy-3-methylglutaryl-Coenzyme A reductase	432	1E-119
A K 0 0 6 5 2 5 Mm. 17834 U.2.16 (YtoM)	NP_612477	hypothetical protein BC000993	523	1E-148
BAB24634.1				
	BAB71062	umamed protein product	521	1E-148
	BAB69025	ALS2CR15	435	1E-122
	AAC50935	islet cell autoantigen p69	271	2E-72
	AAH08640	islet cell autoantigen 1 (69kD)	271	2E-72
	155598	diabetes-associated autoantigen p69	271	2E-72
	NP_071682	islet cell autoantigen 1 isoform 1; islet cell autoantigen 1 (69kD); islet cell autoantigen p69	271	2E-72
	AAA64927	autoantigen p69	268	1E-71
	AAH05922	Similar to islet cell autoantigen 1 (69kD)	366	7E-71
	AAH00993	Unknown (protein for MGC:5250)	234	2E-61
	NP_004959	islet cell autoantigen 1 isoform 2; islet cell autoantigen 1 (69kD); islet cell autoantigen p69	231	2E-60
NM_018779 Mm.103728 U:2.15 (5to19)		CAA06304.1 phosphodiesterase 3A	1379	0
	014432	cGMP-inhibited 3', S'-cyclic phosphodiesterase A (Cyclic GMP inhibited phosphodiesterase A)	1379	0
	,	(CGI-PDEA)		
	NP_000912.2	NP_000912.2 phosphodiesterase 3A, cGMP-inhibited	1379	0
	A44093	cGMP-inhibited cAMP phosphodiesterase (EC 3.1.4), myocardial form	1378	0

٠		312	700	(
	CAA64774.1	64774.1 cyclic nucleotide phosphodiesterase	//9	5
	NP_000913.1	000913.1 phosphodiesterase 3B, cGMP-inhibited	<i>LL</i> 9	0
NM_009350 Mm.8122 U:2.15 (MtoO)	(toO) NP_640336	testis nuclear RNA-binding protein	968	0
NP 033376.1	BAB71416	umamed profess product	968	0
	AAH40229	Similar to testis nuclear RNA-binding protein	892	0
	BAC04125	unnamed protein product	998	0
U 3 6 4 7 5 Mm.1889 U.2.14 (5to11)		009225.1 breast cancer 1, early onset; breast-ovarian cancer, included	1833	5
AAC32323.1	A 54652	hreast/ovarian cancer susceptibility protein BRCA1	1816	0
	FFCPUO GIV	009733 1 hreast cancer 1. early onset isoform BRCA1-delta9-10; breast-ovarian cancer, included	1773	0
	A A B 61673	A A B61673. 1 breast and ovarian cancer susceptibility protein splice variant	1694	0
	NP 009232.	009232.1 breast cancer 1, early onset isoform BRCA1-delta15-17; breast-ovarian cancer, included	1445	0
	NP 009228.	009228.1 breast cancer 1, early onset isoform BRCA1-delta2-10; breast-ovarian cancer, included	1440	0
	NP 009231	009231.1 breast cancer 1, early onset isoform BRCA1-delta14-18; breast-ovarian cancer, included	1420	0
	NP 009230	009230.1 breast cancer 1, early ouset isoform BRCA1-delta14-17; breast-ovarian cancer, included	1419	0
	NP 009234.1	1 breast cancer 1, early onset isoform BRCA1-delta11; breast-ovarian cancer, included	516	1E-144
	NP 009229	009229.1 breast cancer 1, early onset isoform BRCA1-delta9-11; breast-ovarian cancer, included	514	1E-144
	NP 009236.1	1 breast cancer 1, early onset isoform BRCA1-delta9-10-11b; breast-ovarian cancer, included	514	1E-144
	NP_009235	009235.1 breast cancer 1, early onset isoform BRCA1-delta11b; breast-ovarian cancer, included	206	1E-14
NM_011128 Mm.1230 U:2.14 (5to11)		NP_005387.1 pancreatic lipase-related protein 2	748	0
1.00-2000	NP_000927	000927.1 pancreatic lipase	899	0
	pdb/1LPB	Triacylglycerol lipase, panereatic precursor (Pancreatic lipase) (PL	652	0
	1604419A	lipase	646	0

0.

Ŋ

-1

20

 \sim

31

			O. ↑. O.	[100
		NP_006220.1	006220.1 pancreatic lipase-related protein 1	<u>2</u>	1E-180
		AAH25784.1	pancreatic lipase-related protein 1	630	1E-179
		CAA22264.1	A22264.1 dA149D17.1 (PLRP2 (PNLIPRP2, Pancreatic Lipase Related Protein 2 Precursor, EC 3.1.1.3)	217	3E-55
	,		LIKE protein)		
NM_010361 Mm.24118	U:2.14 (5to19)	NP_000845.1	000845.1 glutathione S-transferase theta 2	375	1E-104
NP 034491.1	·				
		AAG02373.1	glutathione S-transferase theta 2	375	1E-104
		AAC13317.1	AAC13317.1 glutathione S-transferase theta 2	364	1E-101
				_	
		XP_056016.1	056016.1 similar to Glutathione S-transferase theta 1 (GST class-theta) (Glutathione transferase T1-1)	239	3E-63
		NP_000844.1	000844.1 glutathione S-transferase theta 1	239	4E-63
		AAH07065.1	glutathione S-transferase theta 1	236	2E-62
NM_007836 Mm.1236	U:2.14 (5to19)	NP_001915.1	001915.1 growth arrest and DNA-damage-inducible, alpha; DNA-damage-inducible transcript 1; DNA	314	2E-84
NP 031862.1			damage-inducible transcript-1; DNA damage-inducible transcript 1		
NM_013607 Mm.3153	U:2.13 (5to19)	NP_002465.1	002465.1 smooth muscle myosin heavy chain 11, isoform SM1	3129	0
NP 038635.1					
		NP_074035.1	074035.1 smooth muscle myosin heavy chain 11, isoform SM2	3083	0
		AAC31665.1	C31665.1 Myosin heavy chain (MHY11) (5'partial)	2907	0
		NP 002464.1	002464.1 myosin, heavy polypeptide 9, non-muscle	2485	0
		P35580	Myosin heavy chain, nonmuscle type B (Cellular myosin heavy chain, type B) (Nonmuscle myosin 2476	ii 247	0
			heavy chain-B) (NMMHC-B)		
		A61231	myosin heavy chain nonmuscle form A	2470	0
		NP_075008.1	075008.1 smooth muscle myosin heavy chain 11, isoform SM3	2180	0
		XP_044702.2	XP_044702.2 similar to Myosin heavy chain, nonmuscle type B (Cellular myosin heavy chain, type B)	1933	3
			(Nomnuscle myosin heavy chain-B) (NMMHC-B		

o ru

		3.1.4	Ì	
	AAA59888.1	59888.1 cellular myosin heavy chain	1916	0
	36971.1	smooth muscle myosin heavy chain	1411	0
	AAH18933.1	AAH18933.1 Unknown (protein for IMAGE:4111094)	1278	0
	1 0 4 2 5 2 4 4 4	Same of the Control o	1258	0
	AAA30349.1	4.30.349.1 Inolumuscie injosui neavy chani (caviaca)		'
	CAA49154.1	smooth muscle mysosin heavy chain	1248	0
	AAA61765.1	461765.1 nonmuscle myosin heavy chain-A	1218	0
	231	myosin heavy chain, nonmuscle, form IIB - human (fragment).	1217	0
	NP_000248.1	000248.1 myosin, heavy polypeptide 7, cardiac muscle, beta	1177	0
	13.1	beta-myosin heavy chain	1176	0
	762	nıyosin alpha heavy chain, cardiac muscle	1176	0
		Myosin heavy chain, cardiac muscle alpha isoform (MyHC-alpha)	1175	0
	CAA79675.1	cardiac alpha-myosin heavy chain	1175	0
	XP_033377.7	033377.7 similar to cardiac alpha-myosin heavy chain	1175	0
	P11055	Myosin heavy chain, fast skeletal muscle, embryonic (Muscle embryonic myosin heavy chain)	1174	0
		(SMHCE)		
	NP_060004.1	060004.1 myosin, heavy polypeptide 2, skeletal muscle, adult	1174	0
	NP_060003.1	060003.1 myosin, heavy polypeptide 4, skeletal muscle	1173	0
	XP_008442.4	008442.4 similar to Myosin heavy chain, skeletal muscle, perinatal (MyHC-perinatal)	1172	0
	138055	myosin heavy chain, perinatal skeletal muscle	1172	0
	NP_002461.1	myosin, heavy polypeptide 3, skeletal muscle, embryonic	1171	0
	NP_002463.1	myosin, heavy polypeptide 8, skeletal muscle, perinatal	1170	0
	CAA37068.1	CAA37068.1 cardiac beta niyosin heavy chain	1165	0
	NP 005954.2	005954.2 myosin, heavy polypeptide 1, skeletal muscle, adult; myosin heavy chain IIx/d	1165	0
	CAC14945.1	C14945.1 dJ756N5.1.1 (Continues in Em:AL133324 as dJ1161H23.3)	1105	0
			500	1
	BAA96036.1	A96036.1 KIAA1512 protein	897	O

10 10 15

				315		ſ
			AAB69327.1	AAB69327.1 smooth muscle myosin heavy chain SM1	824	>
NM_008193 Mm.3624	Mm.3624	U:2.13 (YtoM)	NP_000849	guanylate kinase 1	340	2E-93
NP 032219.1					- 5	000
			AAH09914	guanylate kinase 1	340	2E-93
			106249	guanylate kinase 1	340	2E-93
			AAC50659	guanylate kinase	340	2E-93
			337598	guanylate kinase	340	2E-93
			S68864	guanylate kinase (EC 2.7.4.8) 1	340	2E-93
			Q16774	Guanylate kinase (GMP kinase).	340	2E-93
			8	Similar to guanylate kinase 1	311	6E-85
NM 008542 Mm.27935	Mm.27935	U:2.13(11to19)	NP_005576.2	005576.2 MAID, mothers against decapentaplegic homolog 6; Mothers against decapentaplegic, drosophila,	, 746	0
NP 032568.1				homolog of, 6; MAD (mothers against decapentaplegic, Drosophila) homolog 6		
			AAC00497.1 Smad6	Smad6	746	0
			AACS0792.1 Smad6	Smad6	418	1E-116
			NP_005895.1	005895.1 MAD, mothers against decapentaplegic homolog 7; MAD (mothers against decapentaplegic,	; 298	1E-79
	,			Drosophila) homolog 7; Mothers against decapentaplegic, drosophila, homolog of, 7		
			AAB81354.1		298	1E-79
NM_013891	013891 Min.26768	U:2.13 (MtoO)	AAH21299	prostate epithelium-specific Ets transcription factor	591	1E-168
NP 038919.1					3	2,4
			NP_036523	prostate epithelium-specific Ets transcription factor	291	11:-108
			AAC95296	Ets transcription factor PDEF	591	1E-168
			BAA89543	prostate ets	591	1E-168
NM_011067 Mm.10723	Mm.10723	U:2.12 (YtoM)	NP_058515	period 3; PERIOD, DROSOPHILA, HOMOLOG OF, 3; period circadian protein 3	1116	0
NP 035197.1					1116	C
		·	CAB76084	hypothetical protein	1110	
			BAB32925	period (Drosophila) homolog 3 hPER3	1111	0

10

Ŋ

20

			316		
		P56645 F	Period circadian protein 3 (hP ER3).	1113	0
		AAH26102	Similar to period homolog 3 (Drosophila)	597	1E-170
		02607	period 1; period (Drosophila) homolog 1; hPER; Period, drosophila, homolog of; circadian	563	1E-159
			pacemaker protein RIGUI		
		AAF15544	PERI	563	1E-159
		AAC51765	Rigui	563	1E-159
		534	Period circadian protein 1 (Circadian pacemaker protein Rigui) (hPER).	563	1E-159
		T00018	period protein homolog - human	563	1E-159
		BAC06326	KIAA0482 protein	563	1E-159
		494085	period1.	563	1E-159
		BAA22633	пРет	563	1E-159
		8	KIAA0347 protein	495	1E-139
		NP_073728	period 2 isoform 2; period, Drosophila, homolog of, 2; period circadian protein 2	495	1E-139
			Period circadian protein 2	495	1E-139
		128207	Similar to period homolog 1 (Drosophila)	407	1E-112
		NP_003885	period 2 isoform 1; PERIOD, DROSOPHILA, HOMOLOG OF, 2; period circadian protein 2	206	6E-52
		83709	Per2S	206	6E-52
NM_007377 Mm.6826	U:2.12 (5to11)	BAA31616.2	431616.2 KIAA0641 protein	1406	0
NP 031403.1					
		T00378	KIAA0641 protein	1229	0
		BAB67776.1	KIAA1883 protein	457	1E-127
		XP_055866.4	055866.4 similar to KIAA1883 protein	457	1E-127
		NP_055731,1	055731.1 KIAA1079 protein	414	1E-114
NM_010357 Mm.2662	U:2.11 (5to19)	Q16772	Glutathione S-transferase A3-3 (GST class-alpha)	264	1E-70
NP 034487.1					
		NP_000838.2	glutathione S-transferase A3	263	3E-70
		A49365	glutathione transferase (EC 2.5.1.18) alpha-3 [similarity] - human	261	1E-69
		NP_665683.1	665683.1 glutathione S-transferase A1; GST, class alpha, 1; glutathione S-alkyltransferase A1; glutathione	261	1E-69

Ŋ

			31./ S-aryltransferase A1; S-(hydroxyalkyl)glutathione lyase A1; glutathione S-aralkyltransferase A1;		
			GST-epsilon; glutathione S-transferase 2		
		AAA74634.1	AAA74634.1 glutathione S-transferase A3	261	1E-69
		227110	olutathione transferase (FC 2.5.1.18) A2 - human	259	3E-69
		330	olutathione transferase (FC 2.5.1.18) aloha-2 (clone GTH2) - human	259	4E-69
		1077	CAR07770 1 411571.7 3 (olutathione S-transferase A2)	259	5E-69
		(M) (M)			
		442977	Chain A, Glutathione S-Transferase A1-1 (E.C.2.5.1.18)	259	5E-69
		NP_000837.2	000837.2 glutathione S-transferase A2; glutathione S-transferase 2; GST, class alpha, 2; liver GST2;	258	6E-69
-			glutathione S-alkyltransferase A2; glutathione S-aryltransferase A2; S-(hydroxyalkyl)glutathione		
	· · · · · · · · · · · · · · · · · · ·		lyase A2; glutathione S-aralkyltransferase A2; GST-gamma; HA subunit 2		
		1127144	Chain A, Glutathione Transferase A1-1 Complexed With An Ethacrynic Acid Glutathione	258	1E-68
	1		Conjugate (Mutant R15k)		
		\$20331	glutathione transferase (EC 2.5.1.18) - human	256	2E-68
		DAA00071.1	DAA00071.1 TPA: glutathione transferase A5	256	3E-68
			. (0,110)[]	7.50	t c
		152381	glutathione transferase (EC 2.5.1.18) - human	5 C7	95-08
		XP_167100.2	XP_167100.2 simular to Glutathione S-transferase A1 (GTH1) (HA subunit 1) (GST-epsilon) (GSTA1-1) (GST	253	3E-67
			class-alpha)		
		A56801	glutathione transferase (EC 2.5.1.18) alpha y - human	252	5E-67
		S77958	glutathione transferase (EC 2.5.1.18) alpha-2 (clone GTH2 (+))	248	7E-66
		NP_001503.1	001503.1 glutathione S-transferase A4, glutathione S-alkyltransferase A4; glutathione S-aryltransferase A4,	244	1E-64
			S-(hydroxyalkyl)glutathione lyase A4; glutathione S-aralkyltransferase A4; glutathione transferase		
			A4-4; GST class-alpha; glutathione S-transferase, alpha 4		
NM_008321 Mm.110	U:2.11 (5to11)	NP_002158.1	002158.1 inhibitor of DNA binding 3, dominant negative helix-loop-helix protein; Inhibitor of DNA binding	196	3E-49
NP 032347.1			3, dominant negative, helix-loop-helix		•
		XP_086357.1	XP_086357.1 similar to dJ150O5.2 (Inhibitor of DNA binding 3 (dominant negative helix-loop-helix protein,	195	7E-49
			IR21, HEIR-1))		
, and a second s					

•

)

4(

		002535	DNA-binding protein inhibitor ID-3 (ID-like protein inhibitor HLH 1R21) (Helix-loop-helix	193	3E-48
·		,	protein HEIR-1)		
NM_019393 Mm.116711	11 U:2.1 (YtoM)	CAD44530	polymyositis/scleroderma autoantigen 1	999	0
NP 062266.1					
		CAD56889	polynyositis/scleroderma autoantigen 1	655	0
,		Q06265	Polymyositis/scleroderma autoantigen 1 (Autoantigen PM/Scl 1) (Polymyositis/scleroderma	623	1E-178
			autoantigen 75 kDa) (PM/Scl-75) (P75 polymyositis-scleroderma overlap syndrome associated		-,
	,		autoantigen).		
		NP_005024	polymyositis/scleroderma autoantigen 1, 75kDa; polymyositis/scleroderma autoantigen 1 (75kD)	498	1E-140
		AAA58384	autoantigen	498	1E-140
		G01425	nucleolar 75K autoantigen PM-Scl - human	487	1E-137
		AAA18832	PM-Scl-75 autoantigen	487	1E-137
NM_019670 Mm.28068	8 U:2.09 (MtoO)	BAC03793	unnamed protein product	859	0
NP 062644.1					
		Q9NSV4	Diaphanous protein homolog 3 (Diaphanous-related formin 3) (DRF3).	734	0
		CAC17664	bA218B22.1.1 (novel protein (presumed ortholog of mouse diaphenous-related formin (DIA2))	595	0
		2	(isoform 1))		
		NP_009293	diaphanous 2 isoform 12C	550	1E-156
		CAA75869	DIA-12C protein	550	1E-156
		NP_006720	diaphanous 2 isoform 156	550	1E-156
		CAA75870	DIA-156 protein	550	1E-156
		628090	Diaphanous protein homolog 2 (Diaphanous-related formin 2) (DRF2).	550	1E-156
		NP_112194	diaphanous homolog 3; diaphanous (Drosophila, homolog) 3	526	1Ë-148
		CAB70890	hypothetical protein	526	IE-148
		T46476	hypothetical protein DKFZp434C0931.1	526	1E-148
		NP_005210	diaphanous 1; Diaphanous, Drosophila, homolog of, 1; deafness, autosomal dominant 1;	480	1E-134
			diaphanous (Drosophila, homolog) 1; hDia1		
		AAC05373	diaphanous 1	480	1E-134

S

U) ← i

Ŋ

		319		
	060610	Diaphanous protein homolog 1 (Diaphanous-related formin 1) (DRF1).	480	1E-134
	AAH07411	Unknown (protein for MGC:2554)	368	1E-101
	CAC17665	bA218B22.1.2 (novel protein (presumed ortholog of mouse diaphenous-related formin (DIA2))	332	4E-90
		(translation of cDNA DKFZp434C0931 (Em.AL137718)) (isoform 2))		
	CAB39108	dJ267M20.1 (diaphanous (Drosophila, homolog) 2)	293	2E-78
	BAB14533	unnamed protein product	259	3E-68
	AAH24781	Similar to dishevelled associated activator of morphogenesis 2	206	2E-52
NM_053082 Mm.195498 U:2.09 (YtoM)	NP_003262	transmembrane 4 superfamily member 7; tetraspan TM4SF; novel antigen 2; tetraspanin 4	444	1E-124
NP_444312.1				
	AAH19314	transmembrane 4 superfamily member 7	444	1E-124
	AAH00389	transmembrane 4 superfamily member 7	444	1E-124
	AAC69717	tetraspan TM4SF; Tspan-4	444	1E-124
	AAC51864	tetraspan	444	1E-124
	A59265	tetraspan TSPAN-4 - human.	444	1E-124
	014817	Transmembrane 4 superfamily, member 7 (Novel antigen 2) (NAG-2) (Tetraspanin 4) (Tspan-4).	444	1E-124
	NP_006666	tetraspan NET-5	271	2E-72
	AAC35859	tetraspan NET-5	271	2E-72
	075954	Tetraspan NET-5	27.1	2E-72
NM_009075 Mm.17905 U:2.09 (YtoO)) NP_653164	ribose 5-phosphate isomerase A (ribose 5-phosphate epimerase); RIBOSE 5-PHOSPHATE	450	1E-126
NP 033101.1		ISOMERASE		
	AAH15529	Similar to ribose 5-phosphate isomerase A	430	1E-126
	P49247	Ribose 5-phosphate isomerase (Phosphoriboisomerase).	450	1E-126
N M_022888 Mm.195505 U:2.08 (5to19)) NP_000793	folate receptor 1 (adult)	259	7E-69
	AAA74896	folate-binding protein.	256	4E-68
	NP 000795	T	253	
		7	4	┙

				320		
			CAA49267	folate receptor	253	4E-67
		·	NP_000794	folate receptor 2 precursor	253	5E-67
			AAA17370	folate binding protein	248	1E-65
870056	Mm.3534	U:2.08 (YtoM)	NP_001880	crystallin, zeta; quinone oxidoreductase; NADPH:quinone reductase	519	1E-147
AAB30620.2						
NM_011066 Mm.8471	Mn. 8471	U:2.08 (YtoO)	BAA20804	KIAA0347 protein	1800	0
NP 035196.1						
			NP_073728	period 2 isoform 2; period, Drosophila, homolog of, 2; period circadian protein 2	1791	0
			015055	Period circadian protein 2	1791	0
			NP_002607	period 1; period (Drosophila) homolog 1; hPER; Period, drosophila, homolog of, circadian	ղ 827	0
				pacemaker protein RIGUI		
			AAF15544	PER1	827	0
			AAC51765	Rigui	827	0
)	015534	Period circadian protein 1 (Circadian pacemaker protein Rigui) (hPBR).	827	0
·			T00018	period protein homolog	827	0
			BAC06326.	KIAA0482 protein	827	0
			BAA94085	period1	827	0
			BAA22633	11Per	827	0
			NP_003885	period 2 isoform 1; PERIOD, DROSOPHILA, HOMOLOG OF, 2; period circadian protein 2	595	0
			BAA83709	Per2S	595	0
			BAB32925	period (Drosophila) homolog 3 hPER3	550	1E-155
			P56645	Period circadian protein 3 (IPER3).	550	1E-155
			NP_058515	period 3; PERIOD, DROSOPHILA, HOMOLOG OF, 3; period circadian protein 3	544	1E-153
			CAB76084	hypothetical protein	544	1E-153
			AAH28207	Similar to period homolog 1 (Drosophila) [Homo sapiens].	521	1E-147
			AAH26102.	Similar to period homolog 3 (Drosophila) [Homo sapiens].	340	3E-92

Ŋ

0

15

Ñ

C)

0		0		0.		0	0	0	0	0	0	0		0	0	0	0	0	0	0	0		0	0		0	0	0
1937		1936		1934		1933	1925	1921	1493	1469	1456	1379		1375	1375	1157	1156	1150	1147	1111	1110		1072	1070		1067	1066	1061
321 059145.1 ephrin receptor EphB2 isoform 1 precursor, developmentally-regulated eph-related tyrosine	kinase; elk-related tyrosine kinase; eph tyrosine kinase 3	Ephrin type-B receptor 2 precursor (Tyrosine-protein kinase receptor EPH-3) (DRT) (Receptor	protein-tyrosine kinase HEK5)(ERK).	NP_004433.2 ephrin receptor EphB2 isoform 2 precursor, developmentally-regulated eph-related tyrosine	kinase; elk-related tyrosine kinase; eph tyrosine kinase 3	AAA99310.1 protein-tyrosine kinase	receptor protein-tyrosine kinase	tyrosine kinase precursor	ephrin receptor EphB1 precursor; eph tyrosine kinase 2; ephrin receptor EphB1	AAD02030.1 Eph-like receptor tyrosine kinase hEphB1	Eph-like receptor tyrosine kinase hEphB1b	2004434.2 ephrin receptor EphB3 precursor; human embryo kinase 2; EPH-like tyrosine kinase 2; tyrosine-	protein kinase receptor HEK-2	Eph-like receptor tyrosine kinase hEphB1c	Ephrin type-B receptor 3 precursor (Tyrosine-protein kinase receptor HEK-2).	receptor protein-tyrosine kinase	004429.1 EphA4; Hek8; TYRO1 protein tyrosine kinase; ephrin receptor EphA4	_004431.1 EphA7; Hek11; ephrin receptor EphA7	NP_004430.1 EphA5; Hek7; ephrin receptor EphA5	Ephrin type-A receptor 3 precursor (Tyrosine-protein kinase receptor ETK1) (HEK1) (HEK4).	005224.2 EphA3; Ephrin receptor EphA3 (human embryo kinase 1); eph-like tyrosine kinase 1 (human	embryo kinase 1); ephrin receptor EphA3	Ephrin type-B receptor 4 precursor (Tyrosine-protein kinase receptor HTK).	004435.2 ephrin receptor EphB4 precursor; Ephrin receptor EphB4 (hepatoma transmembrane kinase);	Tyro 11; ephrin receptor EphB4; hepatoma transmembrane kinase	dJ74M1.1.1 (tyrosine kinase isoform 1)	C10351.1 dJ74M1.1.2 (tyrosine kinase isosform 2)	AAA20598.1 tyrosine kinase
NP_059145.		P29323		NP_004433.		AAA99310.1	178842	BAA06506.1	NP_004432.1	AAD02030.	AAD02031.1	NP_004434		AAB94627.1	P54753	178843	NP_004429.	NP_004431.	NP_004430.	P29320	NP_005224.		P54760	NP_004435.		CAC10350.1	CAC10351.	AAA20598.
U:2.08 (YtoM)							٠	٠						-														
Mm.4652																												
L25890	AAA72411.1																											

2

10

<u>___</u>

 $\vec{\sim}$

kinase; tyrosylprotein kinase; tyrosine-protein kinase receptor eek; protein-tyrosine kinase;			
	kinase; tyrosylprotein kinase; tyrosine-protein kinase receptor eek; protein-tyrosine kinase;		
hydroxyaryl-protein kinase			
AAL14195.1 receptor protein tyrosine kinase variant EphB4v1	sphB4v1	686	0
AAH04264.1 Similar to EphB4		946	0
A57174 protein-tyrosine kinase (EC 2.7.1.112) e	rk - human	926	0
NP_004436.1 ephrin receptor EphB6 precursor; tyrosine-protein kinase-defective receptor; ephrin type-B	sine-protein kinase-defective receptor; ephrin type-B	912	0
receptor 6			
AAH37166.1 EphA2		893	°
NP_004422.1 EphA2, ephrin receptor EphA2, epitheli	al cell receptor protein tyrosine kinase	888	0
BAA95983.1 KIAA1459 protein		887	0
AAB94628.1 Eph-like receptor tyrosine kinase hEphE	1d	860	0.
P21709 Ephrin type-A receptor 1 precursor (Tyr	osine-protein kinase receptor EPH	717	0
AAD03058.1 Eph-family protein		713	0
A34076 protein-tyrosine kinase (EC 2.7.1.112)		869	0
BAA03537.1 large erk kinase		969	0
NP_005223.1 BphA1; eph tyrosine kinase 1 erythropoietin-producing hepatoma amplified sequence; oncogene 693	tin-producing hepatoma amplified sequence; oncogene	693	0
EPH; ephrin receptor EphA1); ephtyrosin	EPH; ephrin receptor EphA1); ephtyrosine kinase 1 (erythropoietin-producing hepatoma amplified		
sequence); ephrin receptor EphA1			
AAG43577.1 ephrin receptor EPHA3 secreted form		895	1E-161
CAA81796.1 receptor tyrosine kinase eph		525	1E-148
XP_209519.1 similar to Eph receptor A6 [Mus muscu	[sn	467	1E-131
XP_114973.3 similar to Eph receptor A6 [Mus muscu	[sn	455	1E-127
AAH38796.1 Similar to EphA8		444	1E-124
AAH08655.1 Unknown (protein for IMAGE:3852708		324	4E-88
AAH27940.1 Unknown (protein for MGC:34493)		303	9E-82
CAC19520.1 dJ189K14.1 (cphrin receptor A7)		303	9E-82
XP_209303.1 similar to Ephrin type-A receptor 7 pre	cursor (Tyrosine-protein kinase receptor EHK-3) (Eph	268	3E-71
homology kinase-3) (Receptor protein-tyrosine kinase HEK11)	prosine kinase HEK11)		

			323		
-		NP_775912.1	775912.1 hypothetical protein FLJ33655	700	1B-70
		CAA41565.2	CAA41565.2 tyrosine kinase	228	5E-59
		NP_002101.1	hemopoietic cell kinase	228	5E-59
		TVHUHC	protein-tyrosine kinase (EC 2.7.1.112) hck	228	SE-59
		BAB15482.1	unnamed protein product	228	5E-59
		CAB75606.1	dJ836N17.1 (hemopoietic cell kinase)	228	5E-59
	75	P08631	Tyrosine-protein kinase HCK (p59-HCK/p60-HCK) (Hemopoietic cell kinase).	228	5E-59
		IQCFA	Chain A, Crystal Structure Of Hck In Complex With A Src Family- Selective Tyrosine Kinase	228	5E-59
			Inhibitor		
		NP_005424.1	005424.1 viral oncogene yes-1 homolog 1; proto-oncogene tyrosine-protein kinase YES; Yamaguchi	226	1E-58
			sarcoma oncogene; cellular yes-1 protein		
		BAC04470.1	unnamed protein product	223	1E-57
		NP_722560.1	722560.1 PTK2 protein tyrosine kinase 2 isoform a; focal adhesion kinase 1	223	2E-57
		AAH35404.1	Similar to PTK2 protein tyrosine kinase 2	223	2E-57
		NP_005598.3	NP_005598.3 PTK2 protein tyrosine kinase 2 isoform b; focal adhesion kinase 1	223	2E-57
		NP_002341.1	002341.1 v-yes-1 Yamaguchi sarcoma viral related oncogene homolog; Yamaguchi sarcoma viral (v-yes-1)	222	2E-57
			related oncogene homolog		
		AAH28733.1	Similar to PTK2 protein tyrosine kinase 2	222	2B-57
		AAB50019.1	B50019.1 Lyn B protein	222	2E-57
	· **	NP_002028,1	002028.1 protein-tyrosine kinase fyn isoform a; proto-oncogene tyrosine-protein kinase fyn, src/yes-related	221	4E-57
			novel gene; src-like kinase; c-syn protooncogene; tyrosine kinase p59fyn(T);OKT3-induced		
			calcium influx regulator		
		NP_005347.2	NP_005347.2 lymphocyte-specific protein tyrosine kinase; oncogene LCK; membrane associated protein	221	5E-57
			tyrosine kinase		
		P06239	LCK_HUMAN Proto-oncogene tyrosine-prote	221	5E-57
		AAF34794.1	F34794.1 Proto-oncogene tyrosine-protein kinase LCK (P56-LCK) (LSK) (T cell-specific protein-tyrosine	221	5E-57
		-	kınase).		
		CAA26485.1	CAA26485.1 c-src	220	8E-57
		NP_005408.1	v-src sarcoma (Schmidt-Ruppin A-2) viral oncogene homolog; Protooncogene SRC, Rous	220	8E-57
				ĺ	

Ŋ

7

Ω

		#27C		
		sarcoma; v-src avian sarcoma(Schmidt-Ruppin A-2) viral oncogene homolog		
	1FMK	Crystal Structure Of Human Tyrosine-Protein Kinase C-Src	220	8E-57
	TVHUSC	protein-tyrosine kinase (EC 2.7.1.112) src, neuronal	220	8E-57
	AAA36615.1	src-like tyrosine kinase (put.); putative	219	1E-56
	AAA18225.1	lymphocyte-specific protein tyrosine kinase.	219	2E-56
	AAA59502.1	lymphocyte-specific protein tyrosine kinase.	219	2E-56
	NP_009297.1	009297.1 v-abl Abelson murine leukemia viral oncogene homolog 1 isoform b; Abelson murine leukemia	218	3E-56
		viral (v-abl) oncogene homolog 1		
	NP_005148.1	. 005148.1 v-abl Abelson murine leukemia viral oncogene homolog 1 isoform a; Abelson murine leukemia	218	3E-56
		viral (v-abl) oncogene homolog 1		
,	NP_694592.1	.694592.1 protein-tyrosine kinase fyn isoform b; proto-oncogene tyrosine-protein kinase fyn; src/yes-related	218	4E-56
		novel gene; src-likekinase; c-syn protooncogene; tyrosine kinase p59fyn(T); OKT3-induced		
		calcium influx regulator		`
	NP_002022.1		218	5E-56
	AAC50116.1	Rak	218	5E-56
• .	CAC27542.1	bA702N8.1 (fyn-related kinase)	218	5E-56
	1KSWA	Chain A, Structure Of Human C-Src Tyrosine Kinase (Thr338gly Mutant) In Complex With N6-	218	SE-56
		Benzyl Adp		
	AAB60393.1	proto-oncogene tyrosine-protein kinase.	217	7E-56
	TYHUA	protein-tyrosine kinase (EC 2.7.1.112) abl	217	7E-56
	AAB60394.1	proto-oncogene tyrosine-protein kinase	217	7E-56
	P00519	Proto-oncogene tyrosine-protein kinase ABL1 (p150) (c-ABL).	217	7E-56
	1QPEA	Chain A, Structural Analysis Of The Lymphocyte-Specific Kinase Lck In Complex With Non-	217	9E-56
		Selective And Src Family Selective Kinase Inhibitors.		
	ЗГСК	The Kinase Domain Of Human Lymphocyte Kinase (Lck), Activated Form (Auto-Phosphorylated	217	9E-56
		On Tyr394).		
	AAB33113.2	tyrosine kinase p59fyn(T)	216	2E-55
	1AD5A	Chain A, Src Fanuly Kinase Hck-Amp-Pnp Complex	216	2E-55
	NP_694593.1	694593.1 protein-tyrosine kinase fyn isoform c; proto-oncogene tyrosine-protein kinase fyn; src/yes-related	214	6E-55

			325		
			novel gene; sre-likekinase; c-syn protooncogene; tyrosine kinase p59fyn(T);OKT3-induced		
			calcium influx regulator		
		1QРDА	Chain A, Structural Analysis Of The Lymphocyte-Specific Kinase Lck In Complex With Non-	214	8E-55
			Selective And Src Family Selective Kinase Inhibitors.		
		NP_005237.1	NP_005237.1 fer (fps/fes related) tyrosine kinase (phosphoprotein NCP94); fer (fps/fes related) tyrosine kinase	213	2E-54
		BAB14871.1	unnamed protein	212	3E-54
		CAA28691.1	CAA28691.1 PTK homologous protein (AA 1-507)	210	8E-54
		NP_003206.1	tec protein tyrosine kinase	209	2E-53
		PC1225	protein-tyrosine kinase (EC 2.7.1.112) FAK - human	207	5E-53
		NP_005149.2	NP 005149.2 v-abl Abelson murine leukemia viral oncogene homolog 2 isoforma; Abelson-related protein; arg	207	7E-53
		NP_009298.1	NP_009298.1 v-abl Abelson murine leukemia viral oncogene homolog 2 isoform b; arg; Abelson murine	207	7E-53
			leukemia viral (v-abl) oncogene homolog 2 (arg,		
		AAB60412.1	tyrosine kinase	206	1E-52
		NP_003319.1	TXK tyrosine kinase	206	2E-52
		BAC43747.1	truncated ZAP kinase	205	3E-52
A K 0 1 7 7 5 3 Mm.35505	U:2.08 (YtoO)	NP_689814.1	NP_689814.1 hypothetical protein FLJ38281	312	1E-105
XP 285418	·				
		NP_699189.1	NP_699189.1 hypothetical protein FLJ90396	316	1E-101
		NP_149350.1	NP_149350.1 DKFZP572C163 protein	304	1E-100
		BAC04610.1	BAC04610.1 unnamed protein product	304	1E-99
	,	NP_653290.2	hypothetical protein FLJ32191	300	5E-99
		XP_209968.1	similar to DKFZP572C163 protein	301	8E-99
		T14757	hypothetical protein DKFZp572C163.1	304	2E-98
		AAD23607.1	BC37295_1	309	3E-97
		BAC04309.1	unnamed protein product	317	2E-96
		NP_003427.1	NP_003427.1 zinc finger protein 135 (clone pHZ-17)	315	1E-95
		NP_066358.1	NP_066358.1 zinc finger protein 14 (KOX 6); GIOT-4 for gonadotropin inducible transcription repressor-4	319	5E-95
		BAA86512.1	AA86512.1 KIAA1198 protein	318	5E-95

WO 2005/000335 PCT/US2004/017322

1E-89

AAH36110.1 Similar to zinc finger protein 208

5E-95 3E-94 2E-93 2E-93 3E-93 4E-93 4E-93 8E-93 1E-92 1E-92 3E-92 1E-89 3E-94 3E-94 2E-93 2E-93 1E-92 1E-905E-90 6E-90 1E-90 3E-90 8E-90 1E-91 2E-91 2E-91 2E-91 3E-91 318 303 303 298 275 332 332 280 282 293 293 290 285 301 290 301 295 295 285 278 298 298 294 301 293 294 XP_032674.1 similar to Hypothetical zinc finger protein KIAA1198 Zinc finger protein 16 (Zinc finger protein KOX9). similar to zinc finger protein 347; zinc finger 1111 XP_032678.2 similar to Kruppel-type zinc finger (C2H2) XP_032812.1 similar to hypothetical protein FLJ40981 finger protein 2 placental - human NP_037388.1 zinc finger protein 180 (HHZ168) NP 150630.1 KRAB zinc finger protein KR18 krueppel-type zinc finger protein Similar to zinc finger protein 268 NP 659413.1 hypothetical protein MGC26914 NP_005806.1 Knuppel-type zinc finger (C2H2) NP_110451.1 hypothetical protein FLJ14356 NP_008889.1 zinc finger protein 16 (KOX 9) NP_079009.1 hypothetical protein FLJ14345 NP_085116.1 hypothetical protein FLJ21628 hypothetical protein FLJ40981 hypothetical protein FLJ40981 AAH06528.1 zinc finger protein 43 (HTF6) zinc finger protein 43 (HTF6) NP_003419.1 zinc finger protein 84 (HPF2) 326 BAC04552.1 unnamed protein product zinc finger protein 208 CAD39111.1 hypothetical protein BAB13437.1 KIAA1611 protein CAB94232.2 |zinc finger protein BAB15732.1 FLJ00032 protein ZNF180 XP_030892.2 NP 689815.1 AAH47412.1 AAF71790.1 AAH07307.1 NP 009084.1 NP 003414.1 JE0288 P17020 B32891 Ļ

2 20 20 20 25 25 25 25

	AAH36714.1	H36714.1 Unknown (protein for IMAGE:4846514)	295	1E-89
	T12489	hypothetical protein DKFZp572P0920.1	295	1E-89
	XP_032810.1	032810.1 similar to Zinc finger protein 20 (Zinc finger protein KOX13)(DKFZp572P0920)	295	1E-89
	BAA06541.1	KIAA0065	291	2E-89
	XP_166119.1	XP_166119.1 similar to Zinc finger protein 33A (Zinc finger protein KOX31)HA0946)	291	2E-89
7	NP_008905.1	zinc finger protein 33a; zinc finger and ZAK associated protein with KRAB domain	291	2E-89
	CAC16114.1	bA1021019.1 (zinc finger protein 33a (KOX 31))	291	2E-89
	AAL99923.1	CLL-associated antigen KW 4 splice variant 2	283	2E-89
	NP_003421.1	zinc finger protein 91 (HPF7, HTF10)	290	4E-89
	BAA92587.1	KIAA1349 protein	278	4E-89
	XP_047617.4	similar to Hypothetical zinc finger protein KIAA1349	278	4E-89
	Q9P2J8	Hypothetical zinc finger protein KIAA1349	278	4E-89
	XP_031852.2	similar to Zinc finger protein 84 (Zinc finger protein HPF2)	278	5E-89
	P51814	Zinc finger protein 41	283	1E-88
	NP_700359.1	700359.1 zinc finger protein 41	283	1E-88
	CAC88162.1	bB479F17.3 (zinc finger protein 41)	283	1E-88
	A54661	zinc finger protein ZNF41 - human	283	1E-88
·	NP_115973.1	NP_115973.1 zinc finger protein 347; zinc finger 1111	287	2E-88
	NP_003406.1	zinc finger protein 268	283	5E-88
	AAK69307.1 ZNF268B	ZNF268B	283	5E-88
	AAM28195.1	zinc finger protein 325	285	9E-88
	BAB14183.1	unnamed protein product	285	9E-88
	CAD28491.1	CAD28491.1 hypothetical protein	287	1E-87
	NP_008886.1	NP_008886.1 zinc finger protein 11b (KOX 2)	291	2E-87
	NP_666016.1	zinc finger protein 23; zinc finger protein 32; zinc finger protein 359	281	3E-87
	. P17027	Zinc finger protein 23 (Zinc finger protein KOX16) (DKFZp569D2231).	281	3E-87
	CAD38678.1	CAD38678.1 hypothetical protein	281	3E-87
	AAH15765.1	AAH15765.1 Unknown (protein for MGC:23189)	. 285	3E-87
	NP_065704.1	zinc finger protein 287	290	6E-87

	P35789	Zing finger protein 93 (Zing finger protein HTF34)	787	7F-87
	BAB14401	minima near a minima mental me	201	10 010
	1.10441.000	นเมษามรณ ภาษาแก้ มู่บับนนา	201	0E-0/
	NP_055295.1	zinc finger protein AF020591	288	2E-86
	NP_057528.1	zinc finger protein 226; Kruppel-associated box protein	277	2E-86
	AAF88103.1	zinc finger protein 226	277	2E-86
	91YN9O	Zinc finger protein 226	277	2E-86
	AAF76875.1	zinc finger protein	277	2E-86
	BAC04064.1	unnamed protein product	288	4E-86
	AAH45649.1	Similar to hypothetical protein FLJ32191	270	5E-86
	BAC04764.1	umamed protein product	272	8E-86
		689475.1 hypothetical protein DKFZp571K0837	291	1E-85
	NP_006621.1	zinc finger protein 234; zinc finger protein 269	268	2E-85
	AAF88104.1	ZNF234	268	2E-85
	AAH47570.1	Similar to zinc finger protein 226	277	2E-85
	AAF88107.1	Hypothetical zinc finge	277	2E-85
	NP_004225.2	004225.2 zinc finger protein 93 homolog; zinc finger protein homologous to mouse Zfp93; zinc finger	283	4E-85
		protein homologous to Zfp93 in mouse; zinc finger protein 93 homolog (mouse)		
	Q14588	Zinc finger protein 234 (Zinc finger protein HZF4).	268	4E-85
	137570	zinc finger protein	268	4E-85
	BAC05174.1	unnamed protein product	276	4E-85
	BAB47481.1	KIAA1852 protein	289	5E-85
	AAL58442.1	zinc finger protein 32	289	5E-85
	AAH37209.1	H37209.1 Unknown (protein for MGC:41936)	289	5E-85
	XP_086070.1	086070.1 similar to Zinc finger protein 93 (Zinc finger protein HTF34)	277	7E-85
		060770.2 zinc finger protein 83 (HPF1)	284	7E-85
	XP_209142.1	209142.1 similar to Zinc finger protein 268 (Zinc finger protein HZF3)	273	9E-85
	NP_037512.1	037512.1 zinc finger protein 228	291	1E-84
	AAC51180.1	OS1180.1 kruppel-related zinc finger protein	297	1E-84
	CAD38551.1	O38551.1 hypothetical protein	273	1E-84
-				

. Л

0

15

20

വ

			329		
		XP_032054.2	032054.2 similar to zinc finger protein 28; zinc finger factor X6	273	1E-84
		BAA92634.1	KIAA1396 protein	273	1E-84
NM_023128 Mm.34650	U:2.08 (MtoO)	NP_002570	paralemmin	550	1E-156
NP 075617.1					
		075781	Paralemmin	548	1E-156
		BAA13400	KIAA0270	909	1E-143
		CAA76152	paralemin	451	1E-126
		CAB37401	splice variant	449	1E-126
		T00635	hypothetical protein KIAA0270	429	1E-120
		NP_443749	paralemmin 2	215	2E-55
	_	AAH39306	Similar to paralemmin 2	206	7E-53
		CAC59702	Palm2-AKAP2 fusion protein	204	3E-52
NM_030696 Mm.28632	U:2.08 (5to19)	NP_004198.1	004198.1 solute carrier family 16 (monocarboxylic acid transporters), member 3; monocarboxylate	707	0
NP 109621.1			transporter 3	·	
		NP_037488.1	037488.1 monocarboxylate transporter 3	424	1E-117
		095907	Monocarboxylate transporter 3 (MCT 3)	422	1E-117
		NP_004722.1	004722.1 solute carrier family 16 (monocarboxylic acid transporters), member 7; monocarboxylate	345	1E-93
			transporter 2		
	ſ	AAH30693.1	solute carrier family 16 (monocarboxylic acid transporters), member 7	345	1E-93
		AAC70919.1	AAC70919.1 monocarboxylate transporter 2; MCT2	343	4E-93
		NP_003042.2	003042.2 solute carrier family 16 (monocarboxylic acid transporters), member 1; Solute carrier family 16	311	3E-83
			(monocarboxylic acid transporters),		
		A55568	monocarboxylate transporter 1 - human	311	3E-83
		CAD27707.1	CAD27707.1 monocarboxylate transporter isoform 1	310	5E-83
AB041576 Min.41198	U:2.08 (YtoO)	AAH09942	Unknown (protein for MGC:12595)	254	5E-67
BAA95060.1					
		NP_060629	nudix (nucleoside diphosphate linked moiety X)-type motif 11; hypothetical protein FLJ10628	253	1E-66
				1	-

33(

		IVD OKOOKS	Samuellan de maralia (maralia salah disebutan dan dan dan dan dan dan dan dan dan d		
		ccooon_rv	similiar to mode (increoside diphosphate indeed molety X)-type mont 11; hypothetical protein	251	3E-66
			FLJ10628		
		NP_061967	nudix (nucleoside diphosphate linked moiety X)-type motif 4	236	1E-61
		AAF68858	diphosphoinositol polyphosphate phosphohydrolase type 2 beta	233	1E-60
		AAF68855	diphosphoinositol polyphosphate phosphohydrolase type 2 alpha	229	1E-59
		AAF68857	diphosphoinositol polyphosphate phosphohydrolase type 2 alpha	223	7E-58
NM_019811 Mm.22719	U:2.07 (YtoM)	NP_061147	acetyl-CoA synthetase isoforma; cytoplasmic acetyl-coenzyme A synthetase; acetate-CoA ligase;	1314	0
NP 062785.1			acyl-activating enzyme; acetate thiokinase; acetyl-CoA synthetase		
		AAH12172	Similar to acetyl-CoA synthetase	1312	0
		BAC03849	unnamed protein product	1302	0
*****		NP_644803	acetyl-CoA synthetase isoformb; cytoplasmic acetyl-coenzyme Asynthetase; acetate-CoA ligase;	1137	0
			acyl-activating enzyme, acetate thiokinase; acetyl-CoA synthetase		
		AAH10141	Unknown (protein for MGC:19474)	825	0
		BAB14127	unnamed protein product	824	0
		CAB61786	dI18C9.1.1 (similar to acetyl-coenzyme A synthetase, isoform 1)	701	C
		CAB93422	d11161H23.1 (similar to acetyl-coenzyme A synthetase)	673	0
		AAH39261	Similar to acetyl-Coenzyme A synthetase 2	556	1E-158
		XP_042770	similar to acetyl-CoA synthetase 2 [Mus musculus]	556	1E-158
		AAH44588	similar to acetyl-Coenzyme A synthetase 2 (AMP forming)-like	548	1E-155
		CAC33037	dJ18C9.1.2 (similar to acetyl-coenzyme A synthetase, isoform 2)	525	1E-148
		CAB75500	dJ568C11.3 (novel AMP-binding enzyme similar to acetyl-coenzyme A synthethase (acetate-coA	421	1E-117
			ligase))		
		BAC03853	unnamed protein product	404	1E-112
		BAB47475	KIAA1846 protein	335	2E-91
		NP_078836	hypothetical protein FLJ21963	325	2E-88
		CAC33039	dJ18C9.1.3 (similar to acetyl-coenzyme A synthetase, isoform 3)	218	4E-56
NM 016675 Mm.11706	Mm.117068 U:2.06 (YtoM)	NP 065117	claudin 2	357	1E-98

				331		
NP_057884.1						
NM_011414 Mm.1395	Mm.1395	U:2.05 (YtoO)	NP_003055	secretory leukocyte protease inhibitor precursor; antileukoproteinase; seminal proteinase inhibitor;	182	1E-45
NP 035544.1				mucus proteinase inhibitor		
U89924	Mm.24724	U:2.05 (5to7)	NP_005389.1	005389.1 protein phosphatase 1, regulatory (inhibitor) subunit 5, Phosphatase 1, regulatory inhibitor subunit	545	1E-154
AAB49689.1				5		
			AAD33215.1 PPP1R5	PPPIRS	545	1E-154
			AAM10502.1	AAM10502.1 phosphatase binding 1 protein PTG	909	1E-142
NM_016878 Mm.24680		U:2.04 (YtoM)	NP_036232	aspartyl aminopeptidase	862	0
NP 058574.1						-
	-		AAH00653	aspartyl aminopeptidase	862	0
			BAA92014	unnamed protein product	859	0
			BAA91903	unnamed protein product	721	0
			AAH04854	Similar to aspartyl aminopeptidase	644	0
			BAB13923	unnamed protein product	456	1E-128
			AAH03040	Similar to aspartyl aminopeptidase	281	3E-75
A K 0 0 4 6 3 1 Mm.18939 BAB23425.1	Mm.18939	U:2.04 (5to19)	092871	Phosphomannomutase 1 (PMM 1) (PMMH-22)	511	1E-144
			NP_002667.1	002667.1 phosphomamomutase 1	508	1E-143
			NP_000294.1	_000294.1 phosphomannomutase 2	343	4E-93
NM_023740 Mm.20387	Mn.20387	U:2.04 (5to11)	AAG23766.1 PP3774	PP3774	648	0
NP 076229.1						
			AAH08074.1	AH08074.1 Similar to RIKEN cDNA 1500015N03 gene	638	0
			AAH33157.1	AAH33157.1 similar to Abl-philin 2	523	1E-148
			NP_115703.1	Abl-phiin 2	452	1E-127
NM 010421 Mm.2284	Mm.2284	U:2.04 (YtoM)	NP 000511	hexosaminidase A preproprotein; beta-hexosaminidase alpha chain; beta-N-acetylhexosaminidase;	276	0

			332		
NP_034551.1			N-acetyl-beta-glucosarninidase		
		AAA51827	N-acetyl-alpha-glucosaminidase prepro-polypeptide	068	0
		AAH01138	Similar to hexosaminidase A (alpha polypeptide)	764	0
		AAA51828	N-acetyl-beta-glucosaminidase prepro-polypeptide.	602	1E-172
		168620	beta-hexosaminidase beta-subunit.	602	1E-172
		NP_000512	hexosaminidase B preproprotein; beta-hexosaminidase beta chain; beta-N-acetylhexosaminidase;	602	1E-172
			N-acetyl-beta-glucosaminidase		
-	-				
VM_010444 Mm.119	U:2.04 (MtoO)	NP_002126	nuclear receptor subfamily 4, group A, member 1 isoform a; hormone receptor; growth factor-	936	0
VP_034574.1		ţ	inducible nuclear protein N10; early response protein NAK1; orphan nuclear receptor HMR; TR3		
			orphan receptor; steroid receptor TR3		
,		AAA36763	IR3 orphan receptor	933	0
		NP_006177	nuclear receptor subfamily 4, group A, member 2 isoform a; nur related protein-1 (mouse), human	506	1E-143
			homolog of, transcriptionally inducible nuclear receptor related 1; intermediate-early receptor		
			protein; T-cell nuclear receptor NOT; orphan nuclear receptor NURR1; NGFI-B/nur77 beta-type		
		-	transcription factor homolog		
		NP_775265	nuclear receptor subfamily 4, group A, member 2 isoform d; nur related protein-1 (mouse), human	483	1E-136
	_		homolog of, transcriptionally inducible nuclear receptor related 1; intermediate-early receptor		
			protein; T-cell nuclear receptor NOT; orphan nuclear receptor NURR1; NGFI-B/nur77 beta-type		
			transcription factor homolog		
		AAB33999	NGFI-B/nur77 beta-type transcription factor homolog	478	1E-134
		NP_775263	nuclear receptor subfamily 4, group A, member 2 isoform b; nur related protein-1 (mouse), human	467	1E-131
			homolog of; transcriptionally inducible nuclear receptor related 1; intermediate-early receptor		
			protein; T-cell nuclear receptor NOT; orphan nuclear receptor NURR1; NGFI-B/nur77 beta-type		
			transcription factor homolog		
		092570	Nuclear hormone receptor NOR-1 (Neuron-derived orphan receptor 1) (Mitogen induced nuclear	405	1E-112
			orphan receptor).		
		NP_775292	nuclear receptor subfamily 4, group A, member 3 isoform b; chondrosarcoma, extraskeletal	405	1E-112

			000		
			myxoid, fused to EWS, translocated in extraskeletal chondrosarcoma; neuron derived orphan		
			receptor; mitogen induced nuclear orphan receptor		
		AAB36006	steroid/thyroid orphan receptor homolog gene	404	1E-112
-		NP_008912	nuclear receptor subfamily 4, group A, member 3 isoform a; chondrosarcoma, extraskeletal	404	1E-112
			myxoid, fused to EWS; translocated in extraskeletal chondrosarcoma; neuron derived orphan		<u> </u>
	,		receptor; mitogen induced nuclear orphan receptor		
		BAA11419	neuron derived orphan receptor	401	1E-111
-		AAB02581	mitogen induced nuclear orphan receptor	399	IE-111
		S71930	neuron-derived receptor NOR-1 - human	394	1E-109
	-	NP_775181	nuclear receptor subfamily 4, group A, member 1 isoform b; hormone receptor; growth factor-	393	1E-109
	•		inducible nuclear protein N10; early response protein NAK1; orphan nuclear receptor HMR; TR3		•
			orphan receptor; steroid receptor TR3		
		CAD38550	hypothetical protein	387	1E-107
-		NP_775264	nuclear receptor subfamily 4, group A, member 2 isoforme; nur related protein-1 (mouse), human	306	9E-43
			homolog of, transcriptionally inducible nuclear receptor related 1; intermediate-early receptor		
			protein; T-cell nuclear receptor NOT; orphan nuclear receptor NURR1; NGFI-B/nur77 beta-type		
			transcription factor homolog		
NM_019430 Mm.103723 U.2.04 (YtoM)	2.04 (YtoM)	NP_006530	voltage-dependent calcium channel gamma-3 subunit; neuronal voltage-gated calcium channel	574	IE+164
INF 002303.1			gamma-3 subumt		
		NP_006069	voltage-dependent calcium channel gamma-2 subunit; stargazin; neuronal voltage-gated calcium	433	1E-121
			channel gamma-2 subunit		
		NP_055220	voltage-dependent calcium channel gamma-4 subunit; neuronal voltage-gated calcium channel	316	3E-86
			gamma-4 subunit		
		AAK20031	calcium channel gamma subunit 8	291	9E-79
		NP_114101	voltage-dependent calcium channel gamma-8 subunit, neuronal voltage-gated calcium channel	1 291	9E-79
			gamma-8 subunit		
		Q8WXS5	Voltage-dependent calcium channel gamma-8 subunit (Neuronal voltage-gated calcium channel	1 291	9E-79
			gamma-8 subunit).		<u>_</u>

		AAL50049	voltage-dependent calcium channel gamma-8 subunit	291	1E-78
		AAK15019	putative voltage gated calcium channel gamma-8 subunit CACNG8	227	2E-59
NM_030566 Mm35467	U:2.04 (5to11)	NP_079092.1	079092.1 Fos-related antigen	621	1E-176
NP 085043.1					
NM_011498 Mm.2436	U:2.03 (YtoM)	NP_003661	differentiated embryo chondrocyte expressed gene 1	658	0
NP 035628.1					
		AAK49525	bHLH transcription factor DEC1	652	0
		AAH25968	basic helix-loop-helix domain containing, class B, 3	223	1E-57
		NP_110389	basic helix-loop-helix domain containing, class B, 3; bHLH protein DEC2	223	1E-57
A K 0 1 2 1 6 3 Mm.202683 U.2.03 (7to11)	3 U.2.03 (7to11)	BAA91947.1	A91947.1 unnamed protein product	347	1E-109
BAB28070.1					
		NP_060764.2	060764.2 hypothetical protein FLJ10998	345	1E-109
NM_025703 Mm.182094 U:2.03 (YtoM)	¹⁴ U:2.03 (YtoM)	NP_699164	hypothetical protein MGC45400	150	2E-36
NP 079979.1					
NM_025721 Mm.23402	U:2.02 (5to 19)	NP_663633.1	663633.1 glycosylated 38 kDa sperm protein C-7/8 precursor	221	4E-56
NP 079997.1					
		AAM69364.1	AAM69364.1 glycosylated 38 kDa sperm proiein C-7/8 precursor	216	2E-54
NM_009998 Mm.14177	U:2.02 (11to19) NP		000758.1 cytochrome P450, subfamily IIB (phenobarbital-inducible), polypeptide 6	701	0
11.021120.1		AAF13602.1	cytochrome P450-2B6	697	C
				<u> </u>	,
		AAA52143.1	452143.1 cytochrome P450-IIB	511	1E-144
NM_019692 Mm.42099	U:2.02 (5to 19)	NP_002921.1	002921.1 Ras-like without CAAX 2; Ric-like, expressed in neurons (Drosophila); GTP-binding protein	382	1E-105
NP 062666.1			Roc2		
		AAH18060.1	AAH18060.1 Ric (Drosophila)-like, expressed in neurons	380	1E-104

5

.

7

 \tilde{z}

∠3 ₹)

		_			
		AAB42214.1 rin	rin	380	1E-104
		AAM12636.1	AAM12636.1 Ras family small GTP binding protein RIN	379	1E-104
		AAB64247.1 RIBA	RIBA	281	6E-75
		NP_008843.1	008843.1 Ras-like without CAAX 1; Ric-like, expressed in many tissues (Drosophila); GTP-binding protein	273	2E-72
NM 010368 Mm.3317	U:2,02 (YtoM)	NP 000172	Roct glucironidase, heta	880	
NP_034498.1	,			200	>
	t.	AAH14142	Similar to glucuronidase, beta	984	°
		1BHGA.	Chain A, Human Beta-Glucuronidase At 2.6 A Resolution	362	ľ
NM_009708 Mm.29210 NP 033838.1	U:2.01 (YtoO)	NP_005431	GTP-binding protein Rho7	420	IE-118
		AAH18096	GTP-binding protein Rho7	304	6E-83
		AAB47133	RhoE	284	7E-77
		NP_005159	ras homolog gene family, member E; Rho8; RhoE	284	7E-77
		1M7BA	Chain A, Crystal Structure Of Rnd3RHOE: FUNCTIONAL IMPLICATIONS	259	2E-69
		NP_055285	GTP-binding protein RHO6	235	4E-63
NM_010107 Mm.15675	U:2.01 (5to7)	NP_004419.1	004419.1 ephrin A1 precursor; eph-related receptor tyrosine kinase ligand 1 (tumor necrosis factor, alpha-	353	2E-97
NP 034237.1			induced protein 4)		
		AAH32698.1	H32698.1 ephrin-A1	351	8E-97
NM_012042 Mm.21771	U:2.01 (YtoM)	013616	Cullin homolog 1 (CUL-1).	1508	0
NP 036172.1					
		1LDJA	Chain A, Structure Of The Cul1-Rbx1-Skp1-F Boxskp2 Scf Ubiquitin Ligase Complex	1478	°
		NP_003583	cullin 1	1447	°
		11.DKA	Chain A Structure Of The Cult Dhy 1 Chat E Danda 2 Cell Line 1	000	١

33(

		500		
	AAH34318	18 similar to cullin protein	781	0
	1LDKB	Chain B, Structure Of The Cull-Rbx1-Skp1-F Boxskp2 Scf Ubiquitin Ligase Complex.	687	0
	AAM491	53 cullin 1	535	1E-152
	NP_003582	82 cullin 2	507	1E-143
	AAC51190	90 CUL-2	505	1E-142
	AAC505	45 Hs-CUL-2	479	1E-135
	NP_003581	.81 culin 3	365	1E-100
	AAC36682	82 cullin 3	347	3E-95
	Q93034	Vasopressin-activated calcium-mobilizing receptor (VACM-1) (Cullin homolog 5) (CUL-5).	332	1E-90
	AAB70253	53 vasopressin-activated calcium mobilizing putative receptor protein	329	9E-90
	BAA31670	70 KIAA0695 protein	329	1E-89
	NP_003469	169 Vasopressin-activated calcium-mobilizing receptor-1; Cullin-5 (vasopressin-activated calcium-	n- 329	1E-89
		mobilizing receptor-1)		-
	Q13620	Cullin homolog 4B (CUL-4B).	328	2E-89
	NP_003579	cullin 4B; Cullin-4B	328	2E-89
	AAK16812	12 cullin CUL4B	325	1E-88
	AAC50546	46 Hs-CUL-3.	322	2E-87
	NP_0035	80. cullin 4A	317	98- 3 9
	AAB67315	15 Very similar and perhaps identical to Hs-CUL-4B.; 80-100% similarity to partial sequence	ce 315	2E-85
	,	U58091 (PID:g1381150).		
	BAA33146	46 cullin-4A	276	7E-74
	AAC50547	47 Hs-CUL-4A	228	2E-59
A K 0 1 0 8 2 7 Mm, 22328 U, 2, 01 (YtoM)	(YtoM) NP_079428	128 hypothetical protein FLJ12660	469	1E-132
BAB27209.1	_			
	AAH24919	19 hypothetical protein FLJ12660	466	1E-131
	CAD623	unnamed protein product	252	6E-67
NM 011710 Mm.38433 U:2 (YtoM	oM) P23381	Tryptophanyl-tRNA synthetase (Tryptophan-tRNA ligase) (TRPRS) (IFP53) (hWRS).	860	0

O,

Ŋ

7

Ñ

		33/		
NP_035840.1				
	CAA44450. IFP53	IFP53	858	0
	NP_004175	_004175 tryptophanyl-tRNA synthetase; interferon-induced protein 53	854	0
	CAB94199	B94199 tryptophanyl-tRNA synthetase	554 1E-157	E-157
	CAB94198	B94198 tryptophanyl-tRNA synthetase	226 6E-59	6E-59

Subtable 1C Mixed Genes/Proteins	1C MI	ed Ger	nes/Prote	sui		-
Mouse Gene	Unigene	Unigene Behavior Human		Description	Score	E Value
Protein			Proteins			
		F:-2.07				
		(YtoM)				
AK004731	Mm.19	U:+2.71				
XP_148015	6058	(7to19)	CAA66265	plakophilin 2a	635	0 0
			NP_004563			
				plakophilin 2	614	1e-174
		F:-2.54				
,	·	(YtoO)				
NM_009922	Mm.43	U:+2.55				
NP 034052.1	26	(7to19)	AAH36307	Unknown (protein for IMAGE:5165618)	558	1e-159
			NP_001290	calponin 1, basic, smooth muscle; calponins, basic; Calponin 1	550	16-156
			G02142	smooth muscle cell calponin	548	1e-156
			BAA12983	h1-calponin	510	1e-144
			NP_001830	calponin 3; calponin, acidic	396	1e-110
			NP_004359	calponin 2; Calonin 2	347	7 3e-95
			XP_167021	similar to calponin 2; Calonin 2	266	3 7e-71
			XP_070819	similar to calponin 2; h2-calponin [Mus musculus]	247	7 3e-65
			BAA20887	h2-calponin	228	3 2e-59
		F:-2.11				
		(YtoM)		solute carrier family 7 (cationic amino acid transporter, y+ system), member 9; solute carrier		
NM_021291	Mm.45	U:+3.03		family 7, member 9; solute carrier family 7 (cationic amino acid, transporter, y+ system),		_
NP_067266.1	874	(5to 19)	NP_055085	member 9	754	0
	!					

1 2 2

	CAB54003	glycoprotein-associated amino acid transporter hb0,+AT1	751	0
		solute carrier family 7 (cationic amino acid transporter, y+ system), member 5; Membrane		
	NP_003477	protein E16; Solute carrier family 7, member 5; 4F2 light chain	347	3e-95
	AAC61479.	amino acid transporter E16	347	3e-95
		sodium-independent neutral amino acid transporter LAT1	346	7e-95
	AAH39692	Similar to solute carrier family 7 (cationic amino acid transporter, y+ system), member 5	346	7e-95
	BAA75746	4F2 light chain	346	7e-95
	CAD62619	unnamed protein product	345	2e-94
		Y+L amino acid transporter 1 (y(+)L-type amino acid transporter 1) (y+LAT-1) (Y+LAT1)		
	Q9UM01	(Monocyte amino acid permease 2) (MOP-2).	345	2e-94
		solute carrier family 7, (cationic amino acid transporter, y+ system) member 11;		
	NP_055146	cystine/glutamate transporter	344	2e-94
	NP_003973	solute carrier family 7 (cationic amino acid transporter, y+ system), member 7	343	5e-94
	NP_003974	solute carrier family 7 (cationic amino acid transporter, y+ system), member 6	337	5e-92
		Large neutral amino acids transporter small subunit 2 (L-type amino acid transporter 2)		-
	Q9UHI5	(hLAT2).	328	2e-89
	BAB40574	cystine/glutamate exchanger	328	2e-89
	NP_036376	solute carrier family 7 (cationic amino acid transporter, y+ system), member 8	326	4e-89
	NP 062823	solute carrier family 7, member 10; asc-type amino acid transporter 1	323	4e-88
	BAA95120	y+L amino acid transporter-1	322	1e-87
	AAF05695	L amino acid transporter-2; LAT-2	314	3e-85
	CAD62616	unnamed protein product	210	6e-54
	CAD10393	amino acid transporter	209	1e-53
	NP_620172	amino acid transporter XAT2	207	3e-53

◁	4
~)

] 1e-171			1e-170	1e-169	16-169	7e-88			2e-87	6e-81	8e-81	7 3e-77		7 3e-77	7 3e-77	7 3e-77	3 7e-77	3 4e-76	3 4e-76	3 4e-76	1 2e-75	1 2e-75	1 20 75
				598			265	593	591	322			321	299	299	287		287	287	287	286	283	283	283	281	281	20,4
340				unnamed protein product	keratin 23 isoform a; hyperacetylation-inducible type I keratin; keratin, type I cytoskeletal 23;	cytokeratin 23; type I intermediate filament cytokeratin; histone deacetylase inducible keratin	23	type I intermediate filament cytokeratin	Keratin, type I cytoskeletal 23 (Cytokeratin 23) (K23) (CK 23).	hypothetical protein DKFZp434G032.1	keratin 23 isoform b; hyperacetylation-inducible type I keratin; keratin, type I cytoskeletal 23;	cytokeratin 23; type I intermediate filament cytokeratin; histone deacetylase inducible keratin	23	keratin 20, type I-like, cytoskeletal	similar to Keratin, type I cytoskeletal 20 (Cytokeratin 20) (K20) (CK 20)	Keratin, type I cytoskeletal 19 (Cytokeratin 19) (K19) (CK 19).	keratin 19; keratin, type I cytoskeletal 19; keratin, type I, 40-kd; cytokeratin 19; 40-kDa	keratin intermediate filament precursor gene	unnamed protein product	keratin 17	keratin 19, type I, cytoskeletal	keratin 12 (Meesmann corneal dystrophy); Keratin-12; keratin 12	keratin 15; keratin-15, basic; keratin-15, beta; type I cytoskeletal 15; cytokeratin 15	Keratin, type I cytoskeletal 15 (Cytokeratin 15) (K15) (CK 15).	keratin 13 isoform b; keratin, type I cytoskeletal 13; cytokeratin 13	keratin 13 isoform a; keratin, type I cytoskeletal 13; cytokeratin 13	keratin 13 hma ovtoekalatal long enline form
				BAA92054			NP_056330	AAH28356	Q9C075	T17294		NP_775320		S37780	XP_049979	P08727		NP_002267	BAC04534	NP_000413	KRHU9.	NP 000214	NP_002266	P19012	NP_002265	NP_705694	KRHI 13
	F:-2.05	(YtoO)	U:+2.12	(7to19)					,			,															
			Mm.20	127		,																					
			NM_033373	NP_203537.1									-														

AAA59460	keratin type 16	278	1e-74
NP 005548	keratin 16; keratin, type I cytoskeletal 16; cytokeratin 16	278	2e-74
JC4313	keratin 16, type I, cytoskeletal	278	2e-74
KRHUE	keratin 14, type I, cytoskeletal	277	3e-74
AAH02690	keratin 14 (epidermolysis bullosa simplex, Dowling-Meara, Koebner)	277	3e-74
NP_000517	keratin 14; cytokeratin 14	277	3e-74
NP_003762	type I hair keratin 6; keratin, hair, acidic, 6	264	3e-70
AAH43581	Similar to keratin, hair, acidic, 6	264	3e-70
CAA51914	cytokeratin 20	263	4e-70
NP_002271	type I hair keratin 5; Ha-5; hard keratin, type I, 5	257	3e-68
NP_061889	hypothetical protein FLJ20261	256	6e-68
CAA76387	type I hair keratin 5	256	8e-68
Q92764	Keratin, type I cuticular HA5 (Hair keratin, type I HA5).	256	8e-68
CAA62286	HHa5 hair keratin type I intermediate filament	256	8e-68
XP_039921	similar to keratin 17	253	5e-67
AAH34697	keratin 10 (epidermolytic hyperkeratosis; keratosis palmaris et plantaris)	252	9e-67
P13645	Keratin, type I cytoskeletal 10 (Cytokeratin 10) (K10) (CK 10).	252	9e-67
XP_170564	similar to keratin 17	252	9e-67
NP_004129	lype I hair keratin 3A; Ha-3I; hard keratin, type I,3I; keratin, hair, acidic,3A	251	2e-66
000920	Keratin, type I cuticular HA3-I (Hair keratin, type I HA3-I).	,25i	36-66
KRHU0	keratin 10, type I, cytoskeletal	250	3e-66
NP_002268	type I hair keratin 1; hard keratin, type I, 1; Ha-1; keratin, hair, acidic,1	249	99-99
Q15323	Keratin, type I cuticular HA1 (Hair keratin, type I HA1).	249	7e-66
076011	Keratin, type I cuticular HA4 (Hair keratin, type I HA4).	248	1e-65
NP_002270	type I hair keratin 3B; keratin, hair, acidic,3B; Ha-3II; hard keratin, type I, 3II	248	2e-65
S60034	keratin Ha1, type I, hair	247	4e-65
CAA57956	hair keratin acidic 3-11	246	5e-65
AAH41070	similar to keratin, hair, acidic, 4	246	6e-65

7

7

12

N

Ζ! ₹.)

			7.4.4	-	
		NP_066293	type I hair keratin 4; hard keratin, type I, 4	245	1e-64
		NP 002269	type I hair keratin 2; Ha-2; hard keratin, type I, 2; keratin, hair, acidic,2	245	1e-64
	-	XP 091665	similar to RIKEN cDNA 4733401L19 [Mus musculus]	244	2e-64
		Q14532	Keratin, type I cuticular HA2 (Hair keratin, type I HA2).	244	2e-64
		CAA57-179	hair type I acidic keratin	244	2e-64
		NP 000215	keratin 18	243	4e-64
		CAA82315	cytokeralin 9	243	7e-64
		CAA31377	cytokeratin 18 (424 AA)	243	7e-64
		NP_000217	keratin 9; Keratin-9	243	7e-64
		137459	keratin Ha3-II, type I, hair - human	242	9e-64
		AAH00698	keratin 18	242	1e-63
		AAA59468	keratin-10	239	6e-63
		CAA76389	type I hair keratin 7	236	5e-62
		NP 000412	keratin 10; Keratin-10	236	5e-62
		076015	Keratin, type I cuticular HA8 (Hair keratin, type I HA8).	236	6e-62
		NP_006762	type I hair keratin 8	236	6e-62
		AAH09754	Similar to keratin 18	233	46-61
		NP_003761	type I hair keratin 7	232	9e-61
		BAC03847	unnamed protein product	216	9e-56
	F:-2.87				
	(YtoO)				<u> </u>
X93035 Mm.43	13 U:+2.78				
CAA63603.1 76	(5to19)	AAH08568	Similar to chitinase 3-like 1 (cartilage glycoprotein-39)	537	1e-152
		NP_001267	chitinase 3-like 1; cartilage glycoprotein-39	536	1e-152
		AAH38354	similar to chitinase 3-like 1 (cartilage glycoprotein-39)	535	1e-152
		NP_003456	chitotriosidase; plasma methylumbelliferyl tetra-N-acetylchitotetraoside hydrolase	355	8e-98
		AAG10644	chitotriosidase precursor	355	1e-97

L

1

 \supseteq

C

				0.10		
			1LG1A.	Chain A, Crystal Structure Of Human Chitotriosidase In Complex With Chitobiose	345	8e-95
			NP_003991	chitinase 3-like 2; chondrocyte protein 39	340	3e-93
			Q15782	Chitinase 3-like protein 2 precursor (YKL-39) (Chondrocyte protein 39).	340	3e-93
			AAH11460	chitinase 3-like 2	340	3e-93
			AAB04534	chilinase	340	3e-93
			AAG60019	acidic mammalian chitinase precursor	319	8e-87
			AAO37816	oviductin	274	2e-73
			AAB04126	oviductal glycoprotein	273	5e-73
				oviductal glycoprotein 1, 120kDa (mucin 9, oviductin); mucin 9 (oviductin); oviductal		
			NP_002548	glycoprotein 1, 120kD (mucin 9, oviductin)	273	5e-73
			138605	oviductal glycoprotein	273	5e-73
			NP_068569	eosinophil chemotactic cytokine	226	9e-59
		U:2.04				
		(MtoO)		nuclear receptor subfamily 4, group A, member 1 isoform a; hormone receptor; growth		
NM_010444	Mm.11 F:-2.6	F:-2.6		factor-inducible nuclear protein N10; early response protein NAK1; orphan nuclear receptor		
NP_034574.1	6	(7to11)	NP_002126	HMR; TR3 orphan receptor; steroid receptor TR3	936	0
			AAA36763	TR3 orphan receptor	933	0
······································				nuclear receptor subfamily 4, group A, member 2 isoform a; nur related protein-1 (mouse),		
-				human homolog of; transcriptionally inducible nuclear receptor related 1; intermediate-early		
				receptor protein; T-cell nuclear receptor NOT; orphan nuclear receptor NURR1;		<u> </u>
			NP_006177	NGFI-B/nur77 beta-type transcription factor homolog	506	1e-143
				nuclear receptor subfamily 4, group A, member 2 isoform d; nur related protein-1 (mouse),		
	···			human homolog of; transcriptionally inducible nuclear receptor related 1; intermediate-early		
1	···			receptor protein; T-cell nuclear receptor NOT; orphan nuclear receptor NURR1;		
			NP_775265	NP_775265 NGFI-B/nur77 beta-type transcription factor homolog	483	1e-136
			AAB33999	NGFI-B/nur77 beta-type transcription factor homolog	478	1e-134

1 H J
nuclear receptor subfamily 4, group A, member 2 isoform b; nur related protein-1 (mouse),
human homolog of, transcriptionally inducible nuclear receptor related 1; intermediate-early
receptor protein; T-cell nuclear receptor NOT; orphan nuclear receptor NURR1;
NGFI-B/nur77 beta-type transcription factor homolog
Nuclear hormone receptor NOR-1 (Neuron-derived orphan receptor 1) (Mitogen induced
nuclear orphan receptor).
nuclear receptor subfamily 4, group A, member 3 isoform b; chondrosarcoma, extraskeletal
myxoid, fused to EWS; translocated in extraskeletal chondrosarcoma; neuron derived
orphan receptor; mitogen induced nuclear orphan receptor
steroid/thyroid orphan receptor homolog gene
nuclear receptor subfamily 4, group A, member 3 isoform a; chondrosarcoma, extraskeletal
myxoid, fused to EWS; translocated in extraskeletal chondrosarcoma; neuron derived
orphan receptor; mitogen induced nuclear orphan receptor
neuron derived orphan receptor
mitogen induced nuclear orphan receptor
neuron-derived receptor NOR-1 - human
nuclear receptor subfamily 4, group A, member 1 isoform b; hormone receptor; growth
factor-inducible nuclear protein N10; early response protein NAK1; orphan nuclear receptor
HMR; TR3 orphan receptor; steroid receptor TR3
hypothetical protein
nuclear receptor subfamily 4, group A, member 2 isoform c; nur related protein-1 (mouse),
human homolog of, transcriptionally inducible nuclear receptor related 1; intermediate-early
receptor protein; T-cell nuclear receptor NOT; orphan nuclear receptor NURR1;
NGFI-B/nur77 beta-type transcription factor homolog

2e-97 2e-97 2e-97

1e-113

1e-97 1e-97

1e-161

2e-95 5e-95 7e-95

20

2e-94

		• • • • • •		1172	1171	1163	572	572	512	412	360	360		359	359		359	357	352	352	352	352	352	351	351	349	349	339
345				097818 similar to Probable serine/threonine protein kinase SNF1LK	SNF1-like kinase	gene similar to rat protein kinase (KID2)	KIAA0781 protein	similar to Probable serine/threonine protein kinase SNF1LK	KIAA0781 protein	KIAA0999 protein	Similar to ELKL motif kinase	Ser/Thr protein kinase PAR-1Balpha	MAP/microtubule affinity-regulating kinase 2 isoform b; ELKL motif kinase 1; ELKL motif	kinase	serine/threonine protein kinase	MAP/microtubule affinity-regulating kinase 2 isoform a; ELKL motif kinase 1; ELKL motif	kinase	MAP/microtubule affinity-regulating kinase 1	Cdc25C associated protein kinase C-TAK1	Unknown (protein for MGC:29880)	MARK4 serine/threonine protein kinase	Ser/Thr protein kinase PAR-1A	KIAA1860 protein	probable serine/threonine-specific protein kinase (EC 2.7.1)	MAP/microtubule affinity-regulating kinase 3 long isoform	microtubule affinity-regulating kinase-like1	113605 MAP/microtubule affinity-regulating kinase like 1; MARK4 serine/threonine protein kinase	serine/threonine protein kinase Kp78 splice variant CTAK75a
				XP_097818	NP 775490	BAA95536	BAA34501	XP_041314	BAB91442	BAA76843	AAI-108771	AAK82368.		NP_004945	G01025		NP_059672 kinase	NP_061120	AAC15093	AAH24773	AAL23683	AAK82367	BAB47489	S27966	AAL69982	BAC03375	NP_113605	AAD48007
	U:3.91	(YtoO)	F:-2.39	(11to19)																								
			Mm.23	789																								·
			010831	034961.1																			-	,				

10 10 5

Ø	Ì
4	
m	

				346		
			AAC33487	R31237_1, partial CDS	335	3e-90
			BAA96001	KIAA1477 protein	321	6e-86
			BAA07744	KIAA0096 gene product is related to a protein kinase.	285	5e-75
			P54646	5'-AMP-activated protein kinase, catalytic alpha-2 chain (AMPK alpha-2 chain).	285	6e-75
			AAF86944	HSNFRK	283	2e-74
		U.2.39				
		(YtoO)	-			
NM_010846	Mm.33	F:-2.2		myxovirus resistance protein 1; interferon inducible protein p78; interferon-regulated		
NP_034976.1	966	(5to7)	NP_002453	resistance GTP-binding protein	794	0
			AAA36337	interferon-induced Mx protein	791	0
			BAC04017	unnamed protein product	735	0
		٠		Similar to myxovirus (influenza) resistance 1, homolog of murine (interferon-inducible protein		
			AAH14222	p78)	710	0
	11.27		B33481	interferon-induced viral resistance protein MxB	989	0
			AAA36459	p78-related protein	989	0
			AAC08451	MX2	376	1e-102
			AAC08448	MX2	311	4e-83
			JC4305	dynamin II - human	228	5e-58
٠			P50570	Dynamin 2	226	2e-57
			NP_004936	dynamin 2; Dynamin II	226	2e-57
			B40671	dynamin, internal form 2, short C-terminal form	225	4e-57
		-	AAA02803	dynamin	225	4e-57
			A40671	dynamin, internal form 1, long C-terminal form	223	1e-56
			Q9UQ16	Dynamin 3 (Dynamin, testicular) (T-dynamin).	219	2e-55
			BAA74843	KIAA0820 protein	219	2e-55
			CAB66647	hypothetical protein	217	8e-55
			 -			
			ı			

Ŋ

Г	_	•
7	1	4
ď	•)

				2e-36					3e-62	3e-62	3e-62					2e-87	2e-86	
			· · · ·	150 2	<u> </u>				240 3	240 3	240 3	_				323 2	320 2	_
				15					77	77	2,					3	3,	L
347				hypothetical protein MGC45400					B/B' protein (AA 1-231)	snRNP polypeptide B.	snRNP B' protein					regulator of G protein signalling 16	A28-RGS14p	
				NP_699164_1					CAA33902	AAA60151	CAB57868					AAM12651	AAC16912	
	U:2.03	(YtoM)	F:-2.7	(5to19)		U:2.51	(YtoO)	F:-2.07	(5to11) C)		U:2.23	(YtoM)	F:-3.57	(5to11) /	1	
		_ _	Mm.18					Mm.19							Mm.18	1709		
			NM_025703	NP_079979.1 2094				NM_033174	NP_149409.1 5990						U67189	AAB50619.1		

Table 2

Subtable 2A: Favorable Human Protein Classes

Mouse Gene	Behavior	Human Protein
Mouse Protein		
VM_008341	F:13.28	insulin-like growth factor binding protein 1
NP_032367.1	(5to11)	
		Alternate: Similar to insulin-like growth factor binding protein 1
		Alternate: small IGF-binding-protein
VM 009669	F:8.34 (5to7)	
NP_033799.1	1	amylase
<u> </u>		Subclass: amylase, alpha 2A; pancreatic; Amylase, pancreatic, alpha-2A
		Subclass: amylase, alpha 2B; pancreatic; Amylase, pancreatic, alpha-2B
		Subclass: amylase, alpha 1 A; salivary; Amylase, salivary, alpha-1A
NM 019824	E-5.75 (7to19)	actin related protein 2/3 complex subunit 3; ARP2/3 protein complex subuni
7	1.5.75 (7.015)	
NP_062798.1		p21 Alternate: dJ470L14.3 (novel protein similar to the Arp2/3 protein complex
	:	I '
		subunit p21-Arc (ARC21)) Alternate: similar to ARP 2/3 complex 21 kDa subunit (P21-ARC) (Actin-
		<u> </u>
		related protein 2/3 complex subunit 3)
VM 015763	F:4.93 (5to19)	
- NP 056578.1		Lipin
,		Subclass: Lipin 1
		Subclass: Similar to lipin 1
		Subclass: lipin 2
NM_009117	F·4 72 (5to19)	serum amyloid A1 (SAA1)
	1.4.72 (31015)	,
NP_033143.1		
VM 015805	F:4.48 (5to7)	ATPase IIA
_	ì	
NP 056620 L	1	· · · · · · · · · · · · · · · · · · ·
NP_056620.1		Subclass: ATPase, class 2, member b; ATPase 9B, class II; ATPase 9B, p
NP_056620.1		
NP_056620.1		type Subclass: dJ1114A1.1 (ATPase, class II, type 9A (KIAA0611))
NP 056620.1		type Subclass: dJ1114A1.1 (ATPase, class II, type 9A (KIAA0611)) Subclass: similar to Potential phospholipid-transporting ATPase IIA
NP_056620.1		type Subclass: dJ1114A1.1 (ATPase, class II, type 9A (KIAA0611))
		type Subclass: dJ1114A1.1 (ATPase, class II, type 9A (KIAA0611)) Subclass: similar to Potential phospholipid-transporting ATPase IIA Subclass: Potential phospholipid-transporting ATPase IIB (HUSSY-20)
NM_007706		type Subclass: dJ1114A1.1 (ATPase, class II, type 9A (KIAA0611)) Subclass: similar to Potential phospholipid-transporting ATPase IIA Subclass: Potential phospholipid-transporting ATPase IIB (HUSSY-20) suppressor of cytokine signaling-2; STAT induced STAT inhibitor-2;
		type Subclass: dJ1114A1.1 (ATPase, class II, type 9A (KIAA0611)) Subclass: similar to Potential phospholipid-transporting ATPase IIA Subclass: Potential phospholipid-transporting ATPase IIB (HUSSY-20)
NM_007706 NP_031732.1	F:4.4 (YtoM)	type Subclass: dJ1114A1.1 (ATPase, class II, type 9A (KIAA0611)) Subclass: similar to Potential phospholipid-transporting ATPase IIA Subclass: Potential phospholipid-transporting ATPase IIB (HUSSY-20) suppressor of cytokine signaling-2; STAT induced STAT inhibitor-2; cytokine-inducible SH2 protein 2; (Cish2)
NM_007706 NP_031732.1 NM_008640	F:4.4 (YtoM)	type Subclass: dJ1114A1.1 (ATPase, class II, type 9A (KIAA0611)) Subclass: similar to Potential phospholipid-transporting ATPase IIA Subclass: Potential phospholipid-transporting ATPase IIB (HUSSY-20) suppressor of cytokine signaling-2; STAT induced STAT inhibitor-2; cytokine-inducible SH2 protein 2; (Cish2) lysosomal-associated protein transmembrane 4 alpha; membrane nucleoside
NM_007706 NP_031732.1	F:4.4 (YtoM)	type Subclass: dJ1114A1.1 (ATPase, class II, type 9A (KIAA0611)) Subclass: similar to Potential phospholipid-transporting ATPase IIA Subclass: Potential phospholipid-transporting ATPase IIB (HUSSY-20) suppressor of cytokine signaling-2; STAT induced STAT inhibitor-2; cytokine-inducible SH2 protein 2; (Cish2)
NM_007706 NP_031732.1 NM_008640 NP_032666.1	F:4.4 (YtoM) F:4.09 (5to19)	type Subclass: dJ1114A1.1 (ATPase, class II, type 9A (KIAA0611)) Subclass: similar to Potential phospholipid-transporting ATPase IIA Subclass: Potential phospholipid-transporting ATPase IIB (HUSSY-20) suppressor of cytokine signaling-2; STAT induced STAT inhibitor-2; cytokine-inducible SH2 protein 2; (Cish2) lysosomal-associated protein transmembrane 4 alpha; membrane nucleoside

		Alternate: hypothetical protein DKFZp434J1114.1
M63245	F:3.98 (5to19)	
AAA91867.1		aminolevulinic acid synthase
		Subclass: aminolevulimate, delta-, synthase 1
		Subclass: 5-aminolevu linic acid synthase
		Subclass: 5-aminolevulinic acid synthase, erythroid-specific, mitochor
		precursor (Delta-aminolevulinate synthase) (Delta-ALA synthetase) (A
		E)
		Subclass: aminolevulimate, delta-, synthase 2; Aminolevulinate, delta-
		synthase-2
AK005274	F:3.89 (5to7)	hydroxyacyl glutathione hydrolase; hydroxyacyl glutathione hydrolase
BAB23924.1		glyoxalase 2; Hydroxy acyl glutathione hydrolase; glyoxalase II;
DAD23924.1		
		hydroxyacylglutathione hydroxylase Alternate: similar to HAGH
	<u> </u>	Anemaie: Shillai to FIAGH
NM_026346		
NP 080622.1	F:3.64 (YtoO)	F-box only protein Subclass: F-box only protein 32 isoform 1; muscle atrophy F-box pro
		atrogin-1
		Subclass: F-box only protein 32 isoform 2; muscle atrophy F-box prot
		atrogin-1
		Subclass: F-box only protein 25; F-box protein Fbx25
NM_025298		
-	E-2 45 (X/to)(1)	RNA polymerase III
NP 079574.1	F:3.43 (1 tolv1)	Subclass: RNA polymerase III 80 kDa subunit RPC5
		Dubbluss. 14 (1) polyterorus 12 co les u de les la la la la la la la la la la la la la
NM 022331	F:3.44 (5to19)	homocysteine-inducible, endoplasmic reticulum stress-inducible, ubiq
	1 ' '	l.
_	1	like domain member 1 · MMS-inducible gene
_	<u> </u>	like domain member 1; MMS-inducible gene Alternate: Similar to homocysteine-inducible, endoplasmic reticulum
NP 071726.1		Alternate: Similar to homocysteine-inducible, endoplasmic reticulum
_		like domain member 1; MMS-inducible gene Alternate: Similar to homocysteine-inducible, endoplasmic reticulum inducible, ubiquitin-like domain member 1
_	F:3.41 (YtoO)	Alternate: Similar to homocysteine-inducible, endoplasmic reticulum inducible, ubiquitin-like domain member 1
NP 071726.1 NM_016773	F:3.41 (YtoO)	Alternate: Similar to homocysteine-inducible, endoplasmic reticulum inducible, ubiquitin-like domain member 1
NP 071726.1	F:3.41 (YtoO)	Alternate: Similar to homocysteine-inducible, endoplasmic reticulum inducible, ubiquitin-like domain member 1
NP 071726.1 NM_016773	F:3.41 (YtoO)	Alternate: Similar to homocysteine-inducible, endoplasmic reticulum inducible, ubiquitin-like domain member 1 nucleobindin Subclass: nucleobindin 2 Subclass: nucleobindin 1
NP 071726.1 NM_016773	F:3.41 (YtoO)	Alternate: Similar to homocysteine-inducible, endoplasmic reticulum inducible, ubiquitin-like domain member 1 nucleobindin Subclass: nucleobindin 2
NP 071726.1 NM_016773		Alternate: Similar to homocysteine-inducible, endoplasmic reticulum inducible, ubiquitin-like domain member 1 nucleobindin Subclass: nucleobindin 2 Subclass: nucleobindin 1
NP 071726.1 NM_016773 NP 058053.1 BC017603		Alternate: Similar to homocysteine-inducible, endoplasmic reticulum inducible, ubiquitin-like domain member 1 nucleobindin Subclass: nucleobindin 2 Subclass: nucleobindin 1 Subclass: Nucleobindin 1 precursor (CALNUC)
NP 071726.1 NM_016773 NP 058053.1		Alternate: Similar to homocysteine-inducible, endoplasmic reticulum inducible, ubiquitin-like domain member 1 nucleobindin Subclass: nucleobindin 2 Subclass: nucleobindin 1 Subclass: Nucleobindin 1 precursor (CALNUC) unnamed protein product
NP 071726.1 NM_016773 NP 058053.1 BC017603		Alternate: Similar to homocysteine-inducible, endoplasmic reticulum inducible, ubiquitin-like domain member 1 nucleobindin Subclass: nucleobindin 2 Subclass: nucleobindin 1 Subclass: Nucleobindin 1 precursor (CALNUC) unnamed protein product Alternate: thioredoxin-related transmembrane protein
NP 071726.1 NM_016773 NP 058053.1 BC017603		Alternate: Similar to homocysteine-inducible, endoplasmic reticulum inducible, ubiquitin-like domain member 1 nucleobindin Subclass: nucleobindin 2 Subclass: nucleobindin 1 Subclass: Nucleobindin 1 precursor (CALNUC) unnamed protein product
NP 071726.1 NM_016773 NP 058053.1 BC017603 AAH17603.1	F:3.36 (5to19)	Alternate: Similar to homocysteine-inducible, endoplasmic reticulum inducible, ubiquitin-like domain member 1 nucleobindin Subclass: nucleobindin 2 Subclass: nucleobindin 1 Subclass: Nucleobindin 1 precursor (CALNUC) unnamed protein product Alternate: thioredoxin-related transmembrane protein Alternate: hypothetical protein DKFZp564E1962.1 Alternate: hypothetical protein DJ971N18.2
NP 071726.1 NM_016773 NP 058053.1 BC017603	F:3.36 (5to19) F:3.35 (5to19)	Alternate: Similar to homocysteine-inducible, endoplasmic reticulum inducible, ubiquitin-like domain member 1 nucleobindin Subclass: nucleobindin 2 Subclass: nucleobindin 1 Subclass: Nucleobindin 1 precursor (CALNUC) unnamed protein product Alternate: thioredoxim-related transmembrane protein Alternate: hypothetical protein DKFZp564E1962.1

	1	350
VM 013590	F:3.34 (7to19)	lysozyme precursor
- NP 038618.1		
038018.1		
AK005546	F:3.33 (5to19)	
BAB24114.1		coagulation factor
		Subclass: plasma coagulation factor XI precursor, isoform a; plasma
		thromboplastin antecedent
		Subclass: coagulation factor XI
		Subclass: platelet coagulation factor XI, isoform b; plasma thromboplastin
		antecedent
		Alternate: plasma kallikrein B1 precursor; Kallikrein, plasma; kallikrein 3
		plasma; kallikrein B plasma; Fletcher factor
NM_010286	F:3.32 (5to19)	Glucocorticoid-induced leucine zipper protein (Delta sleep-inducing peptic
NP 034416.1	1	immunoreactor) (DSIP-immunoreactive peptide) (DIP protein) (hDIP) (TS
		22-like protein) (TSC-22R)
		22 INO 5200M) (220 222)
NM_009344	F:3.29 (7to19)	T-cell death associated gene
NP 033370.1	(, , ,	
NF_033370.1		Alternate: pleckstrin homology-like domain, family A, member 1; PQ-rich
		protein
		protein
AK005535	F:3.25 (YtoM)	solute carrier farmily 39 (zinc transporter), member 4
BAB24106.1		
BAB2-1100.1	 	
NM_009864	F:3.24 (YtoO)	Cadherin
NP 033994.1		
		Subclass: cadherin 1, type 1 preproprotein; calcium-dependent adhesion
		protein, epithelia1; cadherin 1, E-cadherin (epithelia1); uvomorulin; cell-
		CAM 120/80; Arc-1
		Subclass: cadherin 2, type 1 preproprotein; N-cadherin 1; cadherin 2, N-
		cadherin (neuronal); neural cadherin; calcium-dependent adhesion protein,
		neuronal Subclass: cadherin 3, type 1 preproprotein; P-cadherin; placental cadherin
		cadherin 3, P-cadherin (placental); calcium-dependent adhesion protein,
		· · · · · · · · · · · · · · · · · · ·
	·	placental Subclass: cadherin 4, type 1 preproprotein; cadherin 4, R-cadherin (retinaI
	,	
	<u> </u>	R-cadherin; retinal cadherin
		Alternate: uvomorulin Alternate: unnamed protein product
·		Panelliare, minarized protein product
NM 007687	F:3.24 (5to7)	
_		cofilin
NP 031713.1		Subclass: cofilin 1 (non-muscle)
		Subclass: cofilin 2 isoform 1
		Subclass: similar to Cofilin, non-muscle isoform (18 kDa

		351
		phosphoprotein)(P18)
		Altermate: destrin (actin depolymerizing factor); destrin
NM_008904	·	-
NP 032930.1	F:3.2 (YtoM)	peroxisome proliferative activated receptor, gamma, coactivator 1
AK005989	F:3.2 (YtoM)	protein disulfide isomerase-related protein
BAB24354.1	:	
DAB2+35+.1		Subclass: protein disulfide isomerase-related protein 5
		
NM 026508	F:3.15 (YtoM)	tumor necrosis factor type 1 receptor associated protein TRAP-1
NP 080784.1		
NF_000704.1		Alternate: Unknown (protein for MGC:15157)
		Alternate: heat shock protein 75
		Alternate: tumor rejection antigen (gp96) 1; Tumor rejection antigen-1
		(gp96) Alternate: unnamed protein product
		Alternate: Unknown (protein for MGC:3823)
		Alternate. Olikhown (protoin for his electry)
NM_021792	F·3 14 (5to 19)	hypothetical protein R30953_1
_	1.5.1 (5.61)	
NP_068564.1		
T. 6. 005 40 4	T.2 11 (5+a11)	ADP-ribosylation factor
NM_025404	F:3.11 (31011)	ADF-Hoosylation factor
NP 079680.1		A LITTLE A D.D. T. L. L. C. A. L. Lille G.
		Subclass: ADP-ribosylation factor 4-like; ADP-ribosylation factor-like 6
		Subclass: similar to ADP-ribosylation factor 4L
		Subclass: ADP-ribosylation factor-like 7 Subclass: ADP-ribosylation factor 4
		Subclass: ADF-Hoosylation factor 4
AK005035	F:3.09 (5to19)	trans ferrin
	1.5.09 (51012)	i i i i i i i i i i i i i i i i i i i
BAB23762.1		
		Alternate: Serum Transferrin
		Alternate: Lactoferrin
	 	Subclass: Lactoferrin (Diferric) Subclass: Lactoferrin (Apo Form)
		Subclass: Lactoferrin (Apo Form) Subclass: Lactoferrin (Copper and Oxalate Form)
		Subclass: Lactolerin (Copper and Oxalate Form) Subclass: neutrophil lactoferrin
		Subclass: Lactotransferrin precursor (Lactoferrin) [Contains: Lactoferron
		}
		A; Lactoferroxin B; Lactoferroxin C]
D. F. 6000000	E-2 00 (5+=10)	CCA_AT/enhancer binding protein (C/EBP), beta; CCAAT/enhancer-bin
NM_009883	F:3.09 (5to19)	1
NP 034013.1	<u> </u>	prote in (C/EBP), beta (transcription factor-5)
·		
NM_021301	1	
NP 067276.1	F:3.08 (YtoM)	solute carrier
		Subclass: solute carrier family 15 (H+/peptide transporter), member 2
		Subclass: solute carrier family 15 (oligopeptide transporter), member 1;
		Human peptide transporter (HPEPT1) mRNA, complete cds
	 	Subclass: Caco-2 oligopeptide transporter
	1	
		stero1/retinol dehydrogenase

		352
NP 038814.1		
		Subclass: 3-hydroxysteroid epimerase; oxidative 3-alpha-hydroxysteroid-
	·	dehydrogenase; 3(alpha->beta)-hydroxysteroid epimerase; retinol
		dehydrogenase; oxidoreductase; NAD+ -dependent 3 alpha-hydroxysteroid
		dehydrogenase
		Subclass: microsomal NAD+-dependent retinol dehydrogenase 4
·		Subclass: orphan short-chain dehydrogenase / reductase; retinol
		dehydrogenase similar protein Subclass: NADP-dependent retinol dehydrogenase/reductase; 3-alpha
		hydroxysteroid dehydrogenase Subclass: 11-cis retinol dehydrogenase (11-cis RDH).
		Subclass: retinol dehydrogenase homolog isoform-1
	·	Subclass: retinol dehydrogenase 5 (11-cis and 9-cis); retinol dehydrogenase
		5 (11-cis and 9-cis)
77. 01.6017	7. 2. 00. (71. 10)	1. 1. (
NM_016917	r:3.08 (/to19)	solute carrier family 11 (proton-coupled divalent metal ion transporters),
NP_058613.1		member 3; ferroportin 1; iron regulated gene 1; ferroportin 1
NM_010004	F:3.08 (5to19)	
NP 034134.1		cytochrome P450, subfamily II
141 054154.1		Subclass: cytochrome P450, subfamily IIC (mephenytoin 4-hydroxylase),
	! !	polypeptide 18; cytochrome P450, subfamily IIC (mephenytoin 4-
		hydroxylase), polypeptide 17; microsomal monooxygenase; flavoprotein-
	į	linked mornooxygenase
		Subclass: cytochrome P450, subfamily IIC (mephenytoin 4-hydroxylase),
		polypeptide 19; mephenytoin 4'-hydroxylase; microsomal monooxygenase;
		xenobiotic monooxygenase; flavoprotein-linked monooxygenase
		Subclass: Cytochrome P450 2C9 (CYPIIC9) (P450 PB-1) (P450 MP-4) (S-
		mephenytoin 4-hydroxylase) (P-450MP).
		Subclass: Cytochrome P450 2C8 (CYPIIC8) (P450 form 1) (P450 MP-
		12/MP-20) (P450 IIC2) (S-mephenytoin 4-hydroxylase)
		Subclass: cytochrome P450 2E1
AB060274		
BAB41208.1	F:3.04 (YtoM)	endothelia1 cell growth factor
		Subclass: endothelial cell growth factor 1 (platelet-derived); thymidine
		phosphorylase; gliostatin
NM_018887	F:3 (7to19)	oxysterol 7alpha-hydroxylase
_	15.5 (71019)	oxysteror 7 arpha-nythoxyrase
NP 061375.1		
NM 024406	F:2.98 (7to19)	fatty acid binding protein 4, adipocyte; A-FABP
- NP 077717.1		
NM_018746	F:2.96 (YtoM)	Inter-alpha-trypsin inhibitor
NP 061216.1		
		Subclass: Inter-alpha-trypsin inhibitor heavy chain H4 precursor (ITI heavy

	•		353
	/	1	chain H4) (Inter-alpha-inhibitor heavy chain 4) (Inter-alpha-trypsin inhibitor
			family heavy chain-related protein) (IHRP) (Plasma kallikrein sensitive
			glycoprotein 120) (PK-120) (GP120) (PRO1851) [Contains: GP57]
			Subclass: pre-alpha (globulin) inhibitor, H3 polypeptide; Inter-alpha
			(globulin) inhibitor, H3 polypeptide
35		1	Subclass: Inter-alpha-trypsin inhibitor heavy chain H1 precursor (ITI heavy
•			chain H1) (Inter-alpha-inhibitor heavy chain 1) (Inter-alpha-trypsin inhibitor
			complex component III) (Serum-derived hyaluronan-associated protein)
			(SHAP)
			Subclass: Inter-alpha-trypsin inhibitor heavy chain H2 precursor (ITI heavy
			chain H2) (Inter-alpha-inhibitor heavy chain 2) (Inter-alpha-trypsin inhibitor
			complex component II) (Serum-derived hyaluronan-associated protein)
			(SHAP)
			Subclass: inter-alpha-trypsin inhibitor family heavy chain-related protein.
	NM 009744	F:2.93 (5to19)	B-cell lymphoma 6 protein; B-cell CLL/lymphoma-6; cys-his2 zinc finger
40	NP_033874.1		transcription factor BCL5; zinc finger protein 51; lymphoma-associated zinc
	1		finger gene on chromosome 3
			Alternate: similar to BcL6-associated zinc finger protein
	NM_017372	F:2.91 (7to19)	lysozyme
4 =	NP 059068.1	 	
45	NM_021313	F:2.91 (5to19)	RING finger protein 25
	NP 067288.1		
	X93035		
50		F:2.87 (YtoO)	chitinase
50	X93035	F:2.87 (YtoO)	Subclass: chitinase 3-like 1; cartilage glycoprotein-39
50	X93035	F:2.87 (YtoO)	Subclass: chitinase 3-like 1; cartilage glycoprotein-39 Subclass: chitotriosidase; plasma methylumbelliferyl tetra-N-
50	X93035	F:2.87 (YtoO)	Subclass: chitinase 3-like 1; cartilage glycoprotein-39 Subclass: chitotriosidase; plasma methylumbelliferyl tetra-N- acetylchitotetraoside hydrolase
50	X93035	F:2.87 (YtoO)	Subclass: chitinase 3-like 1; cartilage glycoprotein-39 Subclass: chitotriosidase; plasma methylumbelliferyl tetra-N- acetylchitotetraoside hydrolase Subclass: chitinase 3-like 2; chondrocyte protein 39
50 .	X93035	F:2.87 (YtoO)	Subclass: chitinase 3-like 1; cartilage glycoprotein-39 Subclass: chitotriosidase; plasma methylumbelliferyl tetra-N- acetylchitotetraoside hydrolase Subclass: chitinase 3-like 2; chondrocyte protein 39 Alternate: oviductal glycoprotein
,	X93035	F:2.87 (YtoO)	Subclass: chitinase 3-like 1; cartilage glycoprotein-39 Subclass: chitotriosidase; plasma methylumbelliferyl tetra-N- acetylchitotetraoside hydrolase Subclass: chitinase 3-like 2; chondrocyte protein 39 Alternate: oviductal glycoprotein Subclass: oviductal glycoprotein 1, 120kDa (mucin 9, oviductin); mucin 9
55	X93035		Subclass: chitinase 3-like 1; cartilage glycoprotein-39 Subclass: chitotriosidase; plasma methylumbelliferyl tetra-N- acetylchitotetraoside hydrolase Subclass: chitinase 3-like 2; chondrocyte protein 39 Alternate: oviductal glycoprotein Subclass: oviductal glycoprotein 1, 120kDa (mucin 9, oviductin); mucin 9 (oviductin); oviductal glycoprotein 1, 120kD (mucin 9, oviductin)
	X93035		Subclass: chitinase 3-like 1; cartilage glycoprotein-39 Subclass: chitotriosidase; plasma methylumbelliferyl tetra-N- acetylchitotetraoside hydrolase Subclass: chitinase 3-like 2; chondrocyte protein 39 Alternate: oviductal glycoprotein Subclass: oviductal glycoprotein 1, 120kDa (mucin 9, oviductin); mucin 9
	X93035 CAA63603.1		Subclass: chitinase 3-like 1; cartilage glycoprotein-39 Subclass: chitotriosidase; plasma methylumbelliferyl tetra-N- acetylchitotetraoside hydrolase Subclass: chitinase 3-like 2; chondrocyte protein 39 Alternate: oviductal glycoprotein Subclass: oviductal glycoprotein 1, 120kDa (mucin 9, oviductin); mucin 9 (oviductin); oviductal glycoprotein 1, 120kD (mucin 9, oviductin)
55	X93035 CAA63603.1 NM_023184 NP_075673.1	F:2.87 (5to11)	Subclass: chitinase 3-like 1; cartilage glycoprotein-39 Subclass: chitotriosidase; plasma methylumbelliferyl tetra-N- acetylchitotetraoside hydrolase Subclass: chitinase 3-like 2; chondrocyte protein 39 Alternate: oviductal glycoprotein Subclass: oviductal glycoprotein 1, 120kDa (mucin 9, oviductin); mucin 9 (oviductin); oviductal glycoprotein 1, 120kD (mucin 9, oviductin) Kruppel-like factor 15; KKLF protein; kidney-enriched Kruppel-like factor
	X93035 CAA63603.1 NM_023184 NP_075673.1 NM_010634	F:2.87 (5to11)	Subclass: chitinase 3-like 1; cartilage glycoprotein-39 Subclass: chitotriosidase; plasma methylumbelliferyl tetra-N- acetylchitotetraoside hydrolase Subclass: chitinase 3-like 2; chondrocyte protein 39 Alternate: oviductal glycoprotein Subclass: oviductal glycoprotein 1, 120kDa (mucin 9, oviductin); mucin 9 (oviductin); oviductal glycoprotein 1, 120kD (mucin 9, oviductin)
55	X93035 CAA63603.1 NM_023184 NP_075673.1	F:2.87 (5to11) F:2.84 (5to19)	Subclass: chitoriosidase; plasma methylumbelliferyl tetra-N- acetylchitotetraoside hydrolase Subclass: chitinase 3-like 2; chondrocyte protein 39 Alternate: oviductal glycoprotein Subclass: oviductal glycoprotein 1, 120kDa (mucin 9, oviductin); mucin 9 (oviductin); oviductal glycoprotein 1, 120kD (mucin 9, oviductin) Kruppel-like factor 15; KKLF protein; kidney-enriched Kruppel-like factor fatty acid binding protein 5 (psoriasis-associated); E-FABP
55	X93035 CAA63603.1 NM_023184 NP_075673.1 NM_010634	F:2.87 (5to11) F:2.84 (5to19)	Subclass: chitotriosidase; plasma methylumbelliferyl tetra-N- acetylchitotetraoside hydrolase Subclass: chitinase 3-like 2; chondrocyte protein 39 Alternate: oviductal glycoprotein Subclass: oviductal glycoprotein 1, 120kDa (mucin 9, oviductin); mucin 9 (oviductin); oviductal glycoprotein 1, 120kD (mucin 9, oviductin) Kruppel-like factor 15; KKLF protein; kidney-enriched Kruppel-like factor fatty acid binding protein 5 (psoriasis-associated); E-FABP Osteoporitin precursor (Bone sialoprotein 1) (Urinary stone protein)
55	NM_023184 NP_075673.1 NM_010634 NP_034764.1	F:2.87 (5to11) F:2.84 (5to19)	Subclass: chitoriosidase; plasma methylumbelliferyl tetra-N- acetylchitotetraoside hydrolase Subclass: chitinase 3-like 2; chondrocyte protein 39 Alternate: oviductal glycoprotein Subclass: oviductal glycoprotein 1, 120kDa (mucin 9, oviductin); mucin 9 (oviductin); oviductal glycoprotein 1, 120kD (mucin 9, oviductin) Kruppel-like factor 15; KKLF protein; kidney-enriched Kruppel-like factor fatty acid binding protein 5 (psoriasis-associated); E-FABP Osteoporatin precursor (Bone sialoprotein 1) (Urinary stone protein) (Secreted phosphoprotein 1) (SPP-1) (Nephropontin) (Uropontin)
55	NM_023184 NP_075673.1 NM_010634 NP_034764.1 NM_009263	F:2.87 (5to11) F:2.84 (5to19)	Subclass: chitinase 3-like 1; cartilage glycoprotein-39 Subclass: chitotriosidase; plasma methylumbelliferyl tetra-N- acetylchitotetraoside hydrolase Subclass: chitinase 3-like 2; chondrocyte protein 39 Alternate: oviductal glycoprotein Subclass: oviductal glycoprotein 1, 120kDa (mucin 9, oviductin); mucin 9 (oviductin); oviductal glycoprotein 1, 120kD (mucin 9, oviductin) Kruppel-like factor 15; KKLF protein; kidney-enriched Kruppel-like factor fatty acid binding protein 5 (psoriasis-associated); E-FABP Osteoporitin precursor (Bone sialoprotein 1) (Urinary stone protein) (Secreted phosphoprotein 1) (SPP-1) (Nephropontin) (Uropontin) Alternate: OPN-a - human (fragment).
55 60	NM_023184 NP_075673.1 NM_010634 NP_034764.1 NM_009263	F:2.87 (5to11) F:2.84 (5to19)	Subclass: chitinase 3-like 1; cartilage glycoprotein-39 Subclass: chitotriosidase; plasma methylumbelliferyl tetra-N- acetylchitotetraoside hydrolase Subclass: chitinase 3-like 2; chondrocyte protein 39 Alternate: oviductal glycoprotein Subclass: oviductal glycoprotein 1, 120kDa (mucin 9, oviductin); mucin 9 (oviductin); oviductal glycoprotein 1, 120kD (mucin 9, oviductin) Kruppel-like factor 15; KKLF protein; kidney-enriched Kruppel-like factor fatty acid binding protein 5 (psoriasis-associated); E-FABP Osteoporatin precursor (Bone sialoprotein 1) (Urinary stone protein) (Secreted phosphoprotein 1) (SPP-1) (Nephropontin) (Uropontin) Alternate: OPN-a - human (fragment). Alternate: OPN-b - human (fragment).
55 60	NM_023184 NP_075673.1 NM_010634 NP_034764.1 NM_009263	F:2.87 (5to11) F:2.84 (5to19)	Subclass: chitinase 3-like 1; cartilage glycoprotein-39 Subclass: chitotriosidase; plasma methylumbelliferyl tetra-N- acetylchitotetraoside hydrolase Subclass: chitinase 3-like 2; chondrocyte protein 39 Alternate: oviductal glycoprotein Subclass: oviductal glycoprotein 1, 120kDa (mucin 9, oviductin); mucin 9 (oviductin); oviductal glycoprotein 1, 120kD (mucin 9, oviductin) Kruppel-like factor 15; KKLF protein; kidney-enriched Kruppel-like factor fatty acid binding protein 5 (psoriasis-associated); E-FABP Osteoporitin precursor (Bone sialoprotein 1) (Urinary stone protein) (Secreted phosphoprotein 1) (SPP-1) (Nephropontin) (Uropontin) Alternate: OPN-a - human (fragment).

		354
NM_007779	F:2.8 (5to19)	colony stimulating factor 1 receptor, formerly McDonough feline sarcoma
NP_031805.1		viral (v-fms) oncogene homolog; C-FMS; Colony-stimulating factor-1
		receptor; oncogene FMS (McDonough feline sarcoma)
,		Alternate: platelet-derived growth factor receptor
		Subclass: platelet-derived growth factor receptor alpha precursor
		Subclass: platelet-derived growth factor receptor beta precursor; beta
		platelet-derived growth factor receptor Subclass: platelet-derived growth factor receptor, beta polypeptide
		Subclass: platelet-derived growth factor receptor, beta polypeptide Subclass: platelet-derived growth factor receptor.
		Alternate: FLT3 receptor tyrosine kinase
		Alternate: fms-related tyrosine kinase 3
		Alternate: protein-tyrosine kinase (EC 2.7.1.112) STK-1 precursor
NM_011825	F:2.8 (5to19)	hypothetical protein FLJ21195 similar to protein related to DAC
NP -035955.1		
NF -033933.1	<u> </u>	Alternate: cysteine knot superfamily 1, BMP antagonist 1; gremlin
AK007707		
BAB25202.1	F:2.79 (Min)	NPD008 protein
BAB23202.1	F.2.79 (WIII)	Alternate: CGI-148 protein
		Titteriate. Cd. 110 ft.
NM_026007	F:2.76 (YtoM)	
NP 080283.1		eukaryotic translation elongation factor
		Subclass: eukaryotic translation elongation factor 1 gamma; elongation
		factor 1-gamma; EF-1-gamma; eEF-1B gamma; translation elongation fac
٠		eEF-1 gamma chain; PRO1608; pancreatic tumor-related protein
		Subclass: Similar to eukaryotic translation elongation factor 1 garmma
		Alternate: pancreatic tumor-related protein
	ļ	Alternate: PRO1608
NR 6 004160	E 2 76 (5+-10)	FK 506 binding protein precursor; FK 506 binding protein 11 (19 kDa)
NM_024169	F:2.76 (5to19)	FR 500 binding protein precinsor, FR 500 binding protein 17 (15 1254)
NP 077131.2		
NM_008061	F:2.75 (5to11)	glucose-6-phosphatase, catalytic
NP_032087.1	<u> </u>	
NM_019806	F:2.74 (5to19)	
NP 062780.1		vesicle-associated membrane protein
		Subclass: VAMP (vesicle-associated membrane protein)-associated prote
		B and C; VAMP-associated protein C; VAMP-associated protein B; VAM
		associated 33 kDa protein
		Subclass: vessicle-associated membrane protein (VAMP)-associated prot
		of 33 kDa; vesicle-associated membrane protein (VAMP), 33 kDa; VAMI
		associated protein A; VAMP (vesicle-associated membrane protein)-
	<u>.</u>	associated protein A (33kD)
NM 022324	F:2.74 (5to19)	
F 1111_02222		
NP_071719.1		stromal cell-derived factor

•		355
	ļ	Subclass: stromal cell-derived factor 2 precursor
		A Iternate: Unknown (protein for MGC:1757)
M12571	F-2 73 (YtoM)	heat shock protein
	1.2.75 (1.014)	Heat shock protein
AAA57234.1	ļ	G. 1. 1 II. (1. 1. 70 LD)
	 	Subclass: Heat shock 70 kDa protein 1 (HSP70.1) (HSP70-1/HSP 70-2) Subclass: heat shock 70kDa protein 1A; heat shock 70kD protein 1A;
		shock-induced protein; dnaK-type molecular chaperone HSP70-1
		Subclass: heat shock 70kDa protein 1-like; Heat-shock 70kD protein-like
		heat shock 70kD protein-like 1; heat shock 70kD protein 1-like
		Subclass: heat shock 70kDa protein 1B; heat shock 70kD protein 1B
		Subclass: heat shock 70kDa protein 2; heat shock 70kD protein 2; Heat-
		shock 70kD protein-2
		Subclass: heat shock 70kDa protein 8 isoform 1; heat shock cognate protein
		71-kDa; heat shock 70kd protein 10; heat shock cognate protein 54;
		constitutive heat shock protein 70; lipopolysaccharide-associated protein
		LPS-associated protein 1
		Subclass: heat shock 70kDa protein 6 (HSP70B')
	7 0 7 0 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1	1
U89415	F:2.73 (5to19)	eukaryotic translation elongation factor 2; polypeptidyl-tRNA
AAC36522.1		
		Alternate: similar to Elongation factor 2 (EF-2)
NM_009242	F:2.73 (5to19)	secreted protein, acidic, cysteine-rich (osteonectin); Osteonectin (secreted
NP 033268.1		protein, acidic, cysteine-rich)
033200.1		Alternate: SPARC-like protein 1 precursor (High endothelial venule prot
		(Hevin) (MAST 9)
		Alternate: Unknown (protein for MGC:45264)
NM_026104	F:2.72 (5to7)	sirnilar to RIKEN cDNA 1700095F04
NP 080380.1		
141 000500.1		
AF294617	F:2.69 (5to7)	6-phosphofructo-2-kinase/fructose-2,6-bisphosphatase
AAG02118.1		
LIGOZIIOIZ		Subclass: 6-phosphofructo-2-kinase/fructose-2,6-biphosphatase 3
		Subclass: inducible 6-phosphofructo-2-kinase/fructose 2,6-bisphosphatas
		Subclass: 6-phosphofructo-2-kinase heart isoform
		Subclass: 6-phosphofructo-2-kinase/fructose-2,6-biphosphatase 4
NM 007952	F-2 65 (5to 10)	glucose regulated protein, 58kDa; glucose regulated protein, 58kD
- .	2.03 (31019)	graduo rogumiou protom, rotori, graduo rogumiou protom, rotor
NP_031978.1		(1) (1) (1) (1) (1) (1) (1) (1) (1) (1)
	<u> </u>	Alternate: protein disulfide-isomerase (EC 5.3.4.1) ER60 prrotein
NM_016674	F:2.65 (5to19)	
NP 057883.1		claudin
.11 007000.1	 	Subclass: claudin 1; senescence-associated epithelial membrane protein 1
		Subclass: claudin 7; Clostridium perfringens enterotoxin receptor–like 2;
,	1	claudin 9
		Subclass: claudin 19

15 -

	_	356
		Subclass: similar to Claudin-19
	50 (1/6, 10)	
NM_008509	F:2.64 (5to19)	lipoprotein lipase
NP 032535.1		·
		Subclass: lipoprotein lipase
		Subclass: Similar to lipoprotein lipase
		Subclass: lipoprotein lipase precursor
		Subclass: endothelial lipase precursor; endothelial cell-derived lipase
		Subclass: lipase C precursor
		Subclass: hepatic lipase precursor
NTM 000600	E-2 62 (7+o1Q)	CD5 antigen-like (scavenger receptor cysteine rich family); Spalpha
_	F.2.03 (7101 9)	CDS antigen-like (seavenger receptor cystome from rammy), epp
NP 033820.1	•	
		Alternate: deleted in malignant brain tumors 1 isoform b precursor
		Alternate: DMBT1/8kb.2 protein
	<u> </u>	Alternate: deleted in malignant brain tumors 1 isoform c precursor
		Alternate: deleted in malignant brain tumors 1 isoform a precursor
1		Alternate: M130 antigen Alternate: M130 antigen precursor, splice form 1
		Alternate: M130 antigen precursor, splice form 3
		Alternate: CD163 antigen; macrophage-associated antigen
		Alternate: scavenger receptor cysteine-rich type 1 protein M160 precurse
		Alternate: scavenger receptor cysteine rich domain containing, group B
ļ		
		domains); scavenger receptor cysteine-rich protein SRCRB-S4D
		Alternate: neurotrypsin precursor; protease, serine, 12; motops in; brain-
		specific serine protease 3; leydin
NM_025459	F:2.63 (7to19)	unnamed protein product
NP 079735.1		
		Alternate: hypothetical protein FLJ20152
X00496	F:2.63 (5to19)	HLA class II histocompatibility antigen, gamma chain (HLA-DR antigen
CAA25191.1		associated invariant chain) (Ia antigen-associated invariant chain) (Ii) (p3
· 	8	(CD74 antigen)
		Alternate: hypothetical protein FLJ13902
		A ALUCANIA DE L'ANDRE
	-	
NM 011435	F:2.61 (5to19)	superoxide dismutase 3, extracellular
-	F:2.61 (5to19)	superoxide dismutase 3, extracellular
NM_011435 NP_035565.1	F:2.61 (5to19)	superoxide dismutase 3, extracellular
NP_035565.1		·
NP 035565.1 NM_007574		superoxide dismutase 3, extracellular complement subcomponent C1q chain C
NP_035565.1		·
NP 035565.1 NM_007574 NP 031600.1		complement subcomponent C1q chain C
NP 035565.1 NM_007574 NP 031600.1	F:2.56 (5to19)	complement subcomponent C1q chain C dynein, axonemal, intermediate polypeptide 1; dynein, axonemal,
NP 035565.1 NM_007574	F:2.56 (5to19)	complement subcomponent C1q chain C
NP 035565.1 NM_007574 NP 031600.1 AK004387 BAC25081.1	F:2.56 (5to19) F:2.55 (YtoO)	complement subcomponent C1q chain C dynein, axonemal, intermediate polypeptide 1; dynein, axonemal, intermediate chain 1; dynein intermediate chain DNAI1
NP_035565.1 NM_007574 NP_031600.1 AK004387	F:2.56 (5to19) F:2.55 (YtoO)	complement subcomponent C1q chain C dynein, axonemal, intermediate polypeptide 1; dynein, axonemal,
NP 035565.1 NM_007574 NP 031600.1 AK004387 BAC25081.1 NM_008330	F:2.56 (5to19) F:2.55 (YtoO)	complement subcomponent C1q chain C dynein, axonemal, intermediate polypeptide 1; dynein, axonemal, intermediate chain 1; dynein intermediate chain DNAI1
NP 035565.1 NM_007574 NP 031600.1 AK004387 BAC25081.1	F:2.56 (5to19) F:2.55 (YtoO)	complement subcomponent C1q chain C dynein, axonemal, intermediate polypeptide 1; dynein, axonemal, intermediate chain 1; dynein intermediate chain DNAI1
NP 035565.1 NM_007574 NP 031600.1 AK004387 BAC25081.1 NM_008330 NP 032087.1	F:2.56 (5to19) F:2.55 (YtoO)	dynein, axonemal, intermediate polypeptide 1; dynein, axonemal, intermediate chain 1; dynein intermediate chain DNAI1 hypothetical protein R30953_1
NP 035565.1 NM_007574 NP 031600.1 AK004387 BAC25081.1 NM_008330 NP 032087.1 NM_009922	F:2.56 (5to19) F:2.55 (YtoO)	dynein, axonemal, intermediate polypeptide 1; dynein, axonemal, intermediate chain 1; dynein intermediate chain DNAI1 hypothetical protein R30953_1

	1		357
			Subclass: calponin 2; Calonin 2
			Subclass: calponin 3; calponin, acidic
	NM_010439	F:2.52 (5to19)	
5	NP 034569.1		High mobility group protein
•			Subclass: high-mobility group box 1; high mobility group box 1; high-
			1
	.	ļ	mobility group (nonhistone chromosomal) protein 1
			Subclass: dJ579F20.1 (high-mobility group (nonhistone chromosomal)
10			protein 1-like 1)
			Subclass: similar to dJ579F20.1 (high-mobility group (nomhistone
			chromosomal) protein 1-like 1
			Subclass: High mobility group protein 1-like 10 (HMG-1L_10)
			Subclass: high-mobility group box 2; high-mobility group (nonhistone
			chromosomal) protein 2
		İ	Subclass: high-mobility group box 3; high-mobility group (nonhistone
		!	chromosomal) protein 4
			Subclass: nonhistone chromosomal protein HMG-2B - hurnan
			Subclass: similar to High mobility group protein 4 (HMG-4) (High mobility
			Company mostoin 20) (HMG 20)
		 	group protein 2a) (HMG-2a) Subclass: HMG2a (high mobility group protein 2a)
7 -		<u> </u>	Subclass: rividza (high mobility group protein 2a) Subclass: similar to HMG2a (high mobility group protein 2a)
15	1		Subclass: SP100-HMG
			Subclass. Sr 100-rivid
	X12905	F:2.51 (5to19)	properdin
	ļ	1.2.51 (51015)	
	CAA31389.1	<u> </u>	
20			
	AK007392	F:2.49 (5to11)	pancreatic elastase
	BAB25008.1		
			Subclass: elastase 1, pancreatic
			Subclass: similar to elastase 1, pancreatic
25			Subclass: pancreatic elastase 2 precursor
			Subclass: pancreatic elastase IIB
1			Subclass: Elastase IIIB precursor (Protease E)
			Subclass: elastase 3, pancreatic (protease E)
		<u> </u>	Subclass: Elastase IIIA precursor (Protease E)
3,0			Subclass: Similar to elastase 3, pancreatic (protease E)
	NM_016847	F:2.48 (5to19)	
	NP 058543.1		arginine vasopressin receptor
			Subclass: arginine vasopressin receptor 1A; V1a vasopressin receptor;
		ľ	vascular/hepatic-type arginine vasopressin receptor; antidiuretic hormone
			vascular/nepatic-type arginine vasopressin receptor, and difference normone
			receptor 1A
35			Subclass: arginine vasopressin receptor 1B; arginine vasopressin receptor 3;
			antidiuretic hormone receptor 1B; vasopressin V1B receptor; pituitary
			i
			vasopressin receptor 3
			Subclass: arginine vasopressin receptor 2
			Alternate: oxytocin receptor
	NM_053177	F:2.47 (7to19)	mucolipidin
4.0	NP 444407		
40	1 111 111 11		

	358				
			Subclass: mucolipin 1		
			Subclass: mucolipin-3		
	NM_023733	F:2.47 (7to19)	carnitine O-octanoyltransferase		
5	NP 076222.1				
			Subclass: carnitine O-octanoyltransferase		
			Subclass: Peroxisomal carnitine octanoyltransferase (COT)		
	AK007588	F:2.47 (5to7)	ring finger protein 7; sensitive to apoptosis gene		
10	XP 135065.1				
	NM_019985	F:2.46 (7to19)	C-type lectin-like receptor-2		
,	NP 064369.1				
			Alternate: Similar to C-type lectin-like receptor-2		
15					
	NM_007572	F:2.45 (5to19)	complement component 1, q subcomponent, alpha polypeptide precursor;		
	NP 031598.1		complement C1q A chain precursor,; complement component C1q, A chain		
	AF218416	F:2.44 (7to19)	tocopherol (alpha) transfer protein (ataxia (Friedreich-like) with vitamin E		
20	AAF25956.1		deficiency); Tocopherol (alpha) transfer protein		
	DA 010642	E-2 41 (X/+-N/)	lichyl-diphosphooligosaccharideprotein glycosyltransferase 63 kDa subunit		
	-	1			
	NP_062616.1		precursor (Ribophorin II) (RPN-II) (RIBIIR)		
			Alternate: Unknown (protein for IMAGE:2961244)		
25			Alternate: Unknown (protein for IMAGE:3532152)		
	NM 010442	E-2 41 (7+o10)	heme oxygenase (decycling)		
-	1 -	1.2.41 (71019)	none oxygenase (decyening)		
	NP 034572.1				
20			Subclass: heme oxygenase (decyclizing) 1 Subclass: heme oxygenase (decyclizing) 2		
30			Subclass, heme daygenase (decyclizing) 2		
	NM 007833	F:2.41 (7to19)	decorin		
	NP 031859.1	``			
	NF 031839.1		Subclass: decorin		
35			Subclass: decorin isoform a preproprotein; dermatan sulphate proteoglycans		
			II; bone proteoglycan II; proteoglycan core protein		
			Subclass: decorin isoform b precursor; dermatan sulphate proteoglycans II;		
		2			
		<u> </u>	bone proteoglycan II; proteoglycan core protein Subclass: decorin isoform c precursor; dermatan sulphate proteoglycans II;		
			I :		
			bone proteoglycan II; proteoglycan core protein		
		1	Alternate: biglycan preproprotein; bone/cartilage proteoglycan-I; dermatan		
			sulphate proteoglycan I		
•			Alternate: asporin (LRR class 1); periodontal ligament as sociated protein 1		
40			The Control of the Table To the		
	NM_016850	F:2.41 (5to19)	Interferon regulatory factor 7 (IRF-7)		
	NP 058546.1				
	-		Subclass: interferon regulatory factor 7 isoform b		
			Subclass: interferon regulatory factor 7 isoform a		
45		 	Subclass: interferon regulatory factor 7 isoform d Subclass: putative interferon regulatory factor 7C.2		
	L	<u> </u>	Dubbiass. parative interferon regulatory factor 70.2		

1	!]
NM_009777	F:2.41 (5to19)	complement component 1, q subcomponent, beta polypeptide precursor;
NP 033907.1		complement component C1q, B chain
141 053707.1		complement compenent cry, a characteristic
NM_008524	F:2.41 (5to19)	lumican
NP 032550.1		
141 052550.1		
NM 010789	F:2.4 (5to19)	
NP 034919.1		Homeobox protein
141_054919.1		Subclass: TALE homeobox protein Meis2d
<u> </u>		Subclass: Homeobox protein Meis2 (Meis1-related protein 1)
		Subclass: Meis1 homolog; Meis1 (mouse) homolog
		Subclass: TALE homeobox protein Meis2b
		Subclass: TALE homeobox protein Meis2a
		Subclass: homeobox protein Meis2 isoform e; Meis (mouse) homolog 2;
		Meis1-related gene 1; TALE homeobox protein Meis2
		Subclass: Homeobox protein Meis3 (Meis1-related protein 2)
		Subclass: similar to Homeobox protein Meis3 (Meis1-related protein 2)
		Alternate: Similar to hypothetical protein DKFZp547H236
		Alternate: Unknown (protein for MGC:2820)
NM 013485	F·2 38 (5to19)	complement protein
-	1.2.50 (5.615)	
NP_038513.1		Subclass: complement component 9
-		Subclass: complement component 8
NM_016906	F:2.37 (5to19)	,
NP 058602.1		sec61 homolog
11 030002.1		Subclass: Sec61 alpha form 1; sec61 homolog
		Subclass: Protein transport protein Sec61 alpha subunit isoform 2 (Sec61
		alpha-2)
	-	Subclass: Similar to Sec61 alpha form 2
		Alternate: Similar to CG9539 gene product
		Alternate: unnamed protein
		Alternate: hypothetical protein
		1 11 - 727771 7271 1200010162
AK004979	F:2.37 (5to19)	similar to RIKEN cDNA 1300010M03
BAB23715.1		
		Alternate: hypothetical protein FLJ20152
		Alternate: unnamed protein product
NM 013922		
HAIAT 012277		
NP_038950.1	F:2.36 (7to19)	zinc finger protein KID3
NM_009369		transforming growth factor, beta-induced, 68kDa; comeal dystrophy; kera
		transforming growth factor, beta-induced, 68kDa; corneal dystrophy; kera epithelin; transforming growth factor, beta-induced, 68kD
NM_009369		transforming growth factor, beta-induced, 68kDa; corneal dystrophy; kera epithelin; transforming growth factor, beta-induced, 68kD Alternate: BIGH3
NM_009369		transforming growth factor, beta-induced, 68kDa; corneal dystrophy; kera epithelin; transforming growth factor, beta-induced, 68kD
NM_009369 NP_033395.1		transforming growth factor, beta-induced, 68kDa; corneal dystrophy; kera epithelin; transforming growth factor, beta-induced, 68kD Alternate: BIGH3
NM_009369 NP_033395.1	F:2.36 (5to11)	transforming growth factor, beta-induced, 68kDa; comeal dystrophy; kera epithelin; transforming growth factor, beta-induced, 68kD Alternate: BIGH3

		360 Subclass: core-binding factor, beta subunit, isoform 1; polyomavirus
	•	
		enhancer binding protein 2, beta subunit; SL3-3 enhancer factor 1 beta
		subunit; SL3/AKV core-binding factor beta subunit
		Subclass: core-binding factor, beta subunit, isoform 2; polyomavirus
		enhancer binding protein 2, beta subunit; SL3-3 enhancer factor 1 beta
		subunit; SL3/AKV core-binding factor beta subunit
AK018585	E-2 35 (5to19)	hypothetical protein FLJ13373
	1.2.23 (3.07)	h)podiousus provincia action is
BAB31292.1		
NM_013594	F:2.35 (5to19)	methyl-CpG binding protein
NP 038622.1	ì	
NF_036022.1		Subclass: methyl-CpG binding protein 1
		Subclass: methyl-CpG binding domain protein 1 isoform 1
		Subclass: methyl-CpG binding domain protein 1 isoform 2
(Subclass: methyl-CpG binding protein splice variant 2
		Subclass: methyl-CpG binding domain protein 1 isoform PCM1
		Subclass: methyl-CpG binding domain protein 1 isoform 3
		Subclass: methyl-CpG binding domain protein 1 isoform 4
Z35168		
CAA84531.1	F:2.34 (YtoM)	collagen
CAA64331.1	F.2.54 (11014)	Subclass: alpha 5 type IV collagen, isoform 2, precurs or; collagen IV, alpha
	1	5 polypeptide; collagen of basement membrane, alpha-5 chain
		Subclass: alpha 5 type IV collagen, isoform 1, precurs or; collagen IV, alpha
·		5 polypeptide; collagen of basement membrane, alpha-5 chain
	1	Subclass: alpha 5 type IV collagen, isoform 3, precurs or; collagen IV, alpha
,		5 polypeptide; collagen of basement membrane, alpha-5 chain
<u> </u>		Subclass: alpha 1 type IV collagen preproprotein; collagen IV, alpha-1
		polypeptide; collagen of basement membrane, alpha-l chain
		Subclass: alpha 2 type IV collagen preproprotein; canstatin
		Subclass: alpha 3 type IV collagen, isoform 1, precursor; collagen IV,
		alpha-3 polypeptide (goodpasture antigen)
	1	Subclass: type IV alpha 6 collagen, isoform B precurs or; collagen IV, alpha
		6 polypeptide; collagen of basement membrane, alpha-6
		Subclass: alpha 4 type IV collagen precursor; Collagen IV, alpha-4
		polypeptide; collagen of basement membrane, alpha-4 chain
	,	
NM_010906		
NP 035036.1	F:2.34 (YtoM)	nuclear factor
		Nuclear factor 1 X-type (Nuclear factor 1/X) (NF1-X) (NFI-X) (NF-I/X)
		(CCAAT-box binding transcription factor) (CTF) (TGGCA-binding protein
		Nuclear factor 1 A-type (Nuclear factor 1/A) (NF1-A) (NFI-A) (NF-I/A)
		(CCAAT-box binding transcription factor) (CTF) (TGGCA-binding protein
		Nuclear factor 1 C-type (Nuclear factor 1/C) (NF1-C) (NFI-C)(NF-I/C)
		(CCAAT-box binding transcription factor) (CTF) (TGGCA-binding protein
		nuclear factor I/B

. 30

	_	361
AK011495	F:2.34 (5to19)	lectin, mannose-binding, 1 precursor; intracellular mannose specific lecti
BAB27655.1		endoplasmic reticulum-golgi intermediate compartment protein 53
NM 010354	F:2.34 (5to19)	gelsolin (amyloidosis, Finnish type); Gelsolin
NP 034484.1	1 ` ′	
NP_034484.1	 	Alternate: Adseverin (Scinderin)
		Alternate: Adseverm (Schiderin) Alternate: villin 1; Villin-1
-	<u> </u>	Alternate: similar to mouse adseverin(D5); similar to PID:g2218019
		Alternate: KIAA1905 protein
		Alternate: Advillin (p92)
		Alternate: Similar to gelsolin (amyloidosis, Finnish type)
		Alternate: Similar to advillin
NM_010238	F:2.33 (7to19)	
NP 034368.1		bromodomain containing protein
11 03 13 00.1		Subclass: bromodomain containing protein 2; female sterile homeotic-rela
		gene 1 (RING3, KIAA9001)
		Subclass: bromodomain containing protein 3; bromodomain-containing 3
		RING3-like gene; open reading frame X
		Subclass: bromodomain-containing protein 4 isoform long; similar to
		RING3; chromosome-associated protein
		Subclass: Similar to bromodomain containing 3
NM 013521	F:2.33 (5to19)	N-formyl peptide receptor
NP 038549.1	1	
NI_038349.1		Subclass: formyl peptide receptor 1
		Subclass: formyl peptide receptor-like 1; lipoxin A4 receptor (formyl pep
	1	
	<u> </u>	receptor related)
		Subclass: formyl peptide receptor-like 2
		Subclass: similar to N-formyl peptide receptor-like 2 protein
		Alternate: orphan G-protein coupled receptor Dez isoform a Alternate: Chemokine receptor-like 1 (G-protein coupled receptor DEZ)
		• • • • • • • • • • • • • • • • • • • •
		protein-coupled receptor ChemR23)
		Alternate: complement component 5 receptor 1 (C5a ligand); complement
		component-5 receptor-2 (C5a ligand)
AK020881		
BAB32239.1	F-2 32 (YtoM)	utrophin; dystrophin-related protein
21 12 2 2 2 2 2 2 2 2 2 2 2 2 2 2 2 2 2		The state of the s
AF320996	F:2.32 (7to19)	
	`	WW domain containing adopter with a solled selfi
AK73808.1		WW domain-containing adapter with a coiled-coil region Subclass: WW domain-containing adapter with a coiled-coil region isofor
		Subclass. w w domain-comaining adapter with a Coned-con region isofor
	<u> </u>	1
		Subclass: WW domain-containing adapter with a coiled-coil region, isofor
		2
		Subclass: WW domain-containing adapter with a coiled-coil region, isofor
	*	2
		Alternative land of the control of t
		Alternate: hypothetical protein PRO1741
		Alternate: bA48B24.1 (A novel protein containing a formin binding prote

		362
		(FBP28) domain)
		Alternate: hypothetical protein
		Alternate: KIAA1844 protein
		Alternate: hypothetical protein MGC1O753.
NM 019830	F:2.32 (7to19)	
NP 062804.1	1 '	- athydronoforaca
NP 002804.1		methyltransferase Subclass: protein arginine N-methyltransferase 1-variant 2
	 	Subclass: Protein arginine N-methyltransferase 1 (Interferon receptor 1-
	İ	1
ļ		bound protein 4)
		Subclass: protein arginine N-methyltransferase 1-variant 1 Subclass: protein arginine N-methyltransferase 1-variant 3
		Subclass: Protein arginine N-methyltransferase 4
		Subclass: Protein arginine N-methyltramsferase 3
	<u> </u>	Subclass: HMT1 hnRNP methyltransfer ase-like 2
		Subclass: HMT1 hnRNP methyltransfer ase-like 3
NM_020564	F:2.32 (5to19)	
_	1.2.32 (3.013)	
NP 065589.1	 	sulfotransferase
		Subclass: hydroxysteroid sulfotransferase SULT2B1a Subclass: sulfotransferase family, cytosolic, 2B, member 1; sulfotransferase
•	. [
		family 2B, member 1
	-	Subclass: hydroxysteroid sulfotransferase SULT2B1b
NM 007614	F:2.31 (YtoM)	catenin (cadherin-associated protein), beta 1, 88kDa; catenin (cadherin-
_		
NP_031640.1		associated protein), beta 1 (88kD); caternin (cadherin-associated protein),
	ļ	beta 1 (88kDa
NM 010501	F:2.3 (YtoM)	
_	, ,	Total Communication of the state of
NP 034631.1	1	Interferon-induced protein Subclass: interferon-induced protein with tetratricopeptide repeats 4
		Subclass: similar to Interferon-induced protein with tetratricopeptide repeats
	1	4 (IFIT-4) (Interferon-induced 60 kDa protein) (IFI-60K) (ISG-60) (CIG49)
		(Retinoic acid-induced gene G protein) (RIG-G)
	-	Subclass: similar to Interferon-induced protein with tetratricopeptide repeats
		2 (IFIT-2) (Interferon-induced 54 kDa protein) (IFI-54K) (ISG-54 K)
		Subclass: interferon-induced protein with tetratricopeptide repeats 1;
		Interferon, alpha-inducible protein (MW 56kD); interferon-induced protein
	· ·	56
		Subclass: retinoic acid- and interferon-inducible protein (58kD)
		Alternate: Unknown (protein for MGC: 14710)
NM_010917	F:2.3 (5to11)	nidogen (enactin); Nidogen; nidogen (en tactin)
NP_035047.1	<u> </u>	
		Subclass: nidogen (enactin)
		Subclass: nidogen 2 (osteonidogen); nidogen 2
	-	Subclass: Similar to nidogen 2 (osteonid ogen)
AK005049	F:2.3 (5to19)	Carboxypeptidase N 83 kDa chain (Carboxypeptidase N regulatory subunit)
ALCO03049	μ.2.2 (21019)	Carooxypophicase is 63 kDa cham (Caro Oxypophicase is regulatory submit)

BAB23775.1	1	363 I
DAD23773.1		Alternate: Similar to RIKEN cDNA 1300018K11 gene
AK009881	F:2.3 (5to19)	endoplasmic reticulum protein 29 precursor; endoplasmic reticulum lu
	F.2.3 (31019)	
BAB26559.1		protein ERp28
NM_009547	F:2.29 (5to7)	zinc finger protein 161 homolog; zinc finger protein homologous to Z
NP 033573.1		in mouse; (Zinc finger protein 5) (hZF5)
NM_018793	F:2.29 (5to19)	
NP 061263.1		tyrosine-protein kinase
		Subclass: IFN-tyk, tyk2=interferon alpha/beta signaling pathway-relat
	1	protein tyrosine kinase [human, Daudi cell line, Peptide Partial, 899 a
		Subclass: similar to Non-receptor tyrosine-protein kinase TYK2
		Subclass: tyrosine kinase 2
		Subclass: janus kinase 1
		Subclass: Janus kinase 2; tyrosine-protein kinase JAK2
I		Subclass: JAK3_HUMAN; JANUS KINASE 3; JAK-3; LEUKOCYT
		JANUS KINASE; L-JAK
<u> </u>		Subclass: JAK3B
NM_018864	F:2.29 (5to19)	
NP 061352.1		Inositol-1(or 4)-monophosphatase (IMPase)
11 001332.1		Subclass: Inositol-1(or 4)-monophosphatase (IMPase) (IMP) (Inositol
	1.	monophosphatase) (Lithium-sensitive myo-inositol monophosphatase
	-	Subclass: inositol(myo)-1(or 4)-monophosphatase 2
		Subclass: similar to Myo-inositol-1 (or 4)-monophosphatase (IMPase)
		(Inositol monophosphatase) (Lithium-sensitive myo-inositol
		monophosphatase A1) Subclass: brain myo-inositol monophosphatase A2b; IMPase A2b
NM_010699	F:2.28 (7to19)	
NP 034829.1	1.2.20 (7.015)	Restate debudre genera
NP 034829.1		lactate dehydrogenase Subclass: lactate dehydrogenase A
		Subclass: Chain A, Human Muscle L-Lactate Dehydrogenase M Chain
		Ternary Complex With Nadh And Oxamate Subclass: lactate dehydrogenase B
	 	Subclass: lactate dehydrogenase C
		Subclass: lactate dehydrogenase A –like
		Subclass: similar to lactate dehydro genase A -like
NM_010187	F:2.28 (7to19)	IgG Fc receptor
_		
NP_034317.1		Es gamma DIIb2
		Fc-gamma-RIIb2 Fc fragment of IgG, low affinity IIb, receptor for (CD32); Fc fragment
		IgG, low affinity II, receptor for (CD32)
		Fc-gamma-RIIb1
NM_029813	F:2.28 (5to19)	zinc finger protein
- NP 084089.1		
11 00-1002.1	<u> </u>	L

	364
	Subclass: zinc finger protein 14 (KOX 6); GIOT-4 for gonadotropin
	inducible transcription repressor-4
	Subclass: similar to zinc finger protein 14 (KOX 6); GIOT-4 for
	gonadotropin inducible transcription repressor-4
	Subclass: similar to zinc finger protein 91 (HPF7, HTF10)
	Subclass: zinc finger protein 180 (HHZ168)
5	Subclass: zinc finger protein 136 (clone pHZ-20)
	Subclass: KIAA1710 protein
	Subclass: similar to Hypothetical zinc finger protein KIAA1710
	Subclass: Zinc finger protein 93 (Zinc finger protein HTF34)
	Subclass: zinc finger protein 135 (clone pHZ-17)
.0	Subclass: zinc finger protein 85 (HPF4, HTF1)
	Subclass: KIAA1198 protein
	Subclass: similar to Hypothetical zinc finger protein KIAA1198
	Subclass: similar to Zinc finger protein 135
	Subclass: similar to Zinc finger protein 93 (Zinc finger protein HTF34)
.5	Subclass: zinc finger protein 91 (HPF7, HTF10)
	Subclass: zinc finger protein 84 (HPF2)
	Subclass: finger protein 2, placental
	similar to KRAB zinc finger protein KR18
	Subclass: zinc finger protein A.F020591
0	Subclass: kruppel-related zinc finger protein
	Subclass: Similar to zinc finger protein 208
	Subclass: zinc finger protein 71; endothelial zinc finger protein induced by
	tumor necrosis factor alpha
	Subclass: zinc finger protein 37 homolog (mouse); Zinc finger protein-37,
	mouse, homolog of; zinc finger protein homologous to Zfp37 in mouse
	Subclass: zinc finger protein 328
. –	Subclass: similar to zinc finger protein 29
25	Subclass: similar to zinc ringer protein 29 Subclass: zinc finger protein 268
	Similar to zinc finger protein 208
	Subclass: Zinc finger protein ZNF45
	Subclass: zinc finger protein 16 (KOX 9)
	Subclass: similar to Zinc finger protein 85
30	Subclass: sintilar to zinc imger protein 65 Subclass: zinc finger protein 43 (HTF6)
	Subclass: similar to Zinc finger protein 35 (Zfp-35)
	Subclass: similar to Zine imger protein 33 (Zip 33) Subclass: zinc finger protein 228
	Subclass: similar to Zinc finger protein 20 (Zinc finger protein KOX13)
	(DKFZp572P0920)
35	Subclass: similar to Zinc finger protein 184
	Subclass: zinc finger protein 177
	Subclass: bB479F17.3 (zinc firager protein 41)
	Subclass: similar to Zinc finger protein 41
	Subclass: zinc finger protein 287
10	Subclass: zinc finger protein 33 1; zinc finger protein 463; C2H2-like zinc
	finger protein
	Subclass: zinc finger protein 27 1
	Subclass: Hypothetical zinc finger protein KIAA1473
	Subclass: riypothetical zinc finger protein KIAA1473 Subclass: similar to Hypothetical zinc finger protein KIAA1473
	Subclass: similar to Hypothetical zinc finger protein KIAA1475 Subclass: similar to Hypothetical zinc finger protein KIAA1956
a	Subclass: Similar to Hypothetical zinc finger protein Subclass: KRAB zinc finger protein
15	
	Subclass: KIAA1956 protein Subclass: TRAF6-inhibitory zinc finger protein; TRAF6-binding zinc finger

	,		365
			protein
			Alternate: hypothetical protein
			Subclass: FLJ40981
50			Subclass: similar to hypothetical protein FLJ40981
			Subclass: hypothetical protein FLJ21628
		<u>.</u>	Subclass: hypothetical protein FLJ32191
		ļ	Subclass: hypothetical protein DKFZp572C163.1
			Subclass: hypothetical protein FLJ30932
55			Subclass: hypothetical protein FLJ14345
		ļ	Subclass: hypothetical protein FLJ90396
			Subclass: hypothetical protein FLJ31526
		<u> </u>	Subclass: hypothetical protein DKFZp572P0920.1
	P. 6.007517	E 0 27 (71-10)	ATID1
60	MM_00/51/	IF:2.27 (7to19)	ancient ubiquitous 46 kDa protein AUP1
	NP_031543.1		
			Alternate: AUP1 homolog
	NM_018816	F:2.27 (5to19)	·
65	NP 061286.1		Apolipoprotein M (ApoM)
.00	141_001200.1		Alternate: similar to Apolip oprotein M (ApoM) (G3a) (HSPC336)
	NM 028740	F:2.27 (5to19)	
	-	1 .2.2	
	NP_083016.1		antichymotrypsin
70			Subclass: alpha1-antichymotrypsin
			Subclass: alpha-1-antichymotrypsin precursor
		 	Subclass: similar to Alpha-1-antichymotrypsin precursor (ACT) Subclass: serine (or cysteine) proteinase inhibitor, clade A (alpha-1
			Subclass: serine (or cysteme) protentase inmotion, clade A (alpha-1
			antiproteinase, antitrypsin), rnember 3
		<u> </u>	
75	NM_016875	1	
•	NP 058571.1	F-2 26 (YtoO)	germ cell specific Y-box binding protein; contrin
	141 050571.1	1:2:20 (1:00)	gorin out opening to the same and the same a
	NM 008362	F:2.26 (5to19)	interleukin 1 receptor, type I precursor; interleukin 1 receptor alpha, type I;
	-	(3.2.2)	
	NP_032388.1	ļ	interleukin receptor 1; antigen CD121a
80	7.5.00005	ļ	
	NM_008295		·
	NP 032321.1	F:2.25 (YtoO)	hydroxysteroid dehydrogenase
			Subclass: hydroxy-delta-5-s teroid dehydrogenase, 3 beta- and steroid delta-
			isomerase 1; Hydroxy-delta-5-steroid dehydrogenase, 3 beta- and steroid
		 	Subclass: 3-beta-hydroxysteroid dehydrogenase/delta-5-delta-4-isomerase.
	· · · · · · · · · · · · · · · · · · ·	<u> </u>	Subclass: hydroxy-delta-5-s-teroid dehydrogenase, 3 beta- and steroid delta-
85		ļ	isomerase 2; Hydroxy-delta-5-steroid dehydrogenase, 3 beta- and steroid
		1	Subclass: dJ871G17.4 (novel 3-beta hydroxysteroid
		1	dehydrogenase/isomerase farmily member)
			Subclass: 3 beta-hydroxy-delta 5-C27-steroid oxidoreductase
	NM 008340	F:2.25 (7to19)	insulin-like growth factor birnding protein, acid labile subunit; INSULIN-
0.0	-		LIKE GROWTH FACTOR BINDING PROTEIN COMPLEX ACID
90	NP_032366.1	1	FIVE OVOM 111 LWCLOK DIMDING LKO LEIM COMILERY WCID
			LABILE CHAIN PRECURSOR
	NM 008343	F:2.25 (5to11)	insulin-like growth factor birding protein 3

10 -

		366
NP 032369.1	[
		Alternate: insulin-like growth factor binding protein 5
NM_008929	F:2.25 (5to19)	interferon-induced double-stranded RNA-activated protein kinase inhibitor
NP 032955.1		
NI 052955.1		
NM 013585	F:2.25 (5to19)	proteasome endop eptidase complex
	,	
NP_038613.1		Subclass: proteasome beta 9 subunit isoform 1 proprotein; proteasome
		1
,		subunit, beta type, 9; proteasome-related gene 2; proteasome chain 7;
	1	macropain chain 7; low molecular mass protein 2; multicatalytic
		endopeptidase cormplex chain 7; proteasome catalytic subunit 1i; proteason
		subunit beta 6i
		Subclass: proteasome beta 9 subunit isoform 2 proprotein; proteasome
		subunit, beta type, 9; proteasome-related gene 2; proteasome chain 7;
		macropain chain 7; low molecular mass protein 2; multicatalytic
•		endopeptidase complex chain 7; proteasome catalytic subunit 1i; proteason
		subunit beta 6i
NM_008035	F-2 24 (VtoO)	folate-binding protein
	1.2.24 (1.00)	lolate-oniting proton
NP_032061.1		
		Subclass: folate binding protein 2
		Subclass: folate receptor 3 precursor
	<u> </u>	Subclass: folate receptor 1 (adult) Subclass: similar to Folate receptor gamma precursor (FR-gamma) (Folate
	· ·	1
		receptor 3)
	<u> </u>	gene predicted from cDNA with a complete coding sequence; caught by
NM_025649		gene predicted from cDNA with a complete coding sequence, caught by
NP 079925.1	F:2.24 (Min)	MAD Two 2
NM_011656	F:2.24	tuftelin 1
NP 035786.1	(11to19)	
		Alternate: Similar to tuftelin 1
		Alternate: unnamed protein product
NM_021099		
NP 066922.1	F-2 23 (YtoO)	protein kinase tran smembrane receptor
141 000322.1	1.2.25 (1.00)	Subclass: KIT protein
		Subclass: colony stimulating factor receptor
		Subclass: platelet-derived growth factor receptor
		Subclass: FLT3 receptor tyrosine kinase
		Subclass: vascular endothelial growth factor receptor
		Subclass: fibroblast growth factor receptor
		Subclass: ret proto-oncogene
NM_008290		
NP 032316.1	F:2.23 (YtoO)	hydroxysteroid dehydrogenase
	1	Subclass: hydroxysteroid (17-beta) dehydrogenase 2
		Subclass: 11beta-hydroxysteroid dehydrogenase (EC 1.1.1.146) type 2
······		

NM_008180	F:2.23 (5to19)	glutathione synthetase
 NP 032206.1		·
INF 032200.1		
NM 007468	F:2.22 (7to11)	
_	1.2.22 ()	
NP_031494.1		apolipoprote in
		Subclass: apolipoprotein A-IV
		Subclass: Apolipoprotein A-IV precursor (Apo-AIV)
NTM 007490	F:2.22 (7to11)	
NM_007489	(۱۱۵۱۱) کئی۔1.2	
NP 031515.1		receptor nuclear translocator/transcription factor
		Subclass: aryl hydrocarbon receptor nuclear translocator-like
	,	Subclass: aryl hydrocarbon receptor nuclear translocator; Arnt
	•	Subclass: Aryl hydrocarbon receptor nuclear translocator 2 (ARNT pro
		2)
	 	Subclass: PAS protein 3
······		Subclass: BMAL1 protein (Brain and muscle ARNT-like 1) (Member of
		PAS protein 3) (Basic-helix-loop-helix-PAS orphan MOP3) (BHLH-PA
,		PAS protein 3) (Basic-henx-loop-henx-PAS orphan MOI 3) (BitEll-1 P
		protein JAP3)
		Subclass: transcription factor BMAL2
		Subclass: brain-muscle-ARNT-like transcription factor 2a
		Subclass: brain-muscle-ARNT-like transcription factor 2b
		Subclass: brain-muscle-ARNT-like transcription factor 2c
		Subclass: brain-muscle-ARNT-like transcription factor 2d
		Subclass: bHLH-PAS transcription factor MOP9
		Subclass: bHLH-PAS transcription factor MOP9
		Subclass: Similar to transcription factor BMAL2
		Subclass: brain and muscle Ah receptor nuclear translocator-like protein
	Ì	BMAL1e
		Subclass: brain and muscle Ah receptor nuclear translocator-like protei
		•
		BMAL1d
NM_022985	F-2 22 (7to 10)	protein associated with PRK1; hypothetical protein; associated with PRI
1/1/1_022983	F.2.22 (71019)	protein associated with river, hypothetical protein, associated with river
NP 075361.2		
		Alternate: hypothetical protein
		Alternate: zinc finger protein 216
		Alternate: similar to protein associated with PRK1; hypothetical protein
		associated with PRK1
NM_018754	F:2.22 (5to7)	stratifin
NP 061224.1		
NP_001224.1		Alternate: Similar to stratifin
		Alternate: tyrosine 3/tryptophan 5 -monooxygenase activation protein
		Subclass: tyrosine 3/tryptophan 5 -monooxygenase activation protein, z
	-	
		polypeptide; protein kinase C inhibitor protein-1; phospholipase A2
		Subclass: tyrosine 3/tryptophan 5 -monooxygenase activation protein, t
		polypeptide; 14-3-3 protein tau
	 	Subclass: tyrosine 3-monooxygenase/tryptophan 5-monooxygenase
		activation protein, beta polypeptide; 14-3-3 protein beta/alpha; Protein

		368
]		Subclass: tyrosine 3-monooxygenase/tryptophan 5-monooxygenase
		activation protein, gamma polypeptide Subclass: tyrosine 3/tryptophan 5 -monooxygenase activation protein, eta
		polypeptide
NM_007825	F:2.22 (5to19)	cytochrome P45O, subfamily VIIB, polypeptide 1; oxysterol 7alpha-
NP_031851.1		hydroxylase
		Alternate: Cytochrome P450 7A1 (Cholesterol 7-alpha-monooxygenase)
-		(CYPVII) (Cholesterol 7-alpha-hydroxylase)
NM_021354	F:2.21 (7to19)	
NP_067329.1		developmentally regulated GTP binding protein
		Subclass: developmentally regulated GTP binding protein 2
		Subclass: developmentally regulated GTP binding protein 1; neural
	ì	precursor cell expressed, developmentally down-regulated 3;
		developmentally regulated GTP-binding protein 1
NM_007912	F-2 21 (5to19)	Epidermal growth factor receptor
	(, ,	
NP_031938.1		Subclass: epidermal growth factor receptor (erythroblastic leukemia viral (v
		erb-b) oncogene homolog, avian); epidermal growth factor receptor (aviam
		erythroblastic leukemia viral (v-erb-b) oncogene homolog); Epidermal
		growth factor receptor
		Subclass: p110 epidermal growth factor receptor
		Subclass: v-erb-a erythroblastic leukemia viral oncogene homolog 4; avian
		erythroblastic leukemia viral (v-erb-b2) oncogene homolog 4; v-erb-a avian
		erythroblastic leukemia viral oncogene homolog-like 4
	ł	Subclass: v-erb-b2 erythroblastic leukemia viral oncogene homolog 3
		(avian); Transformation gene ERBB-3; v-erb-b2 avian erythroblastic
		leukemia viral oncogene homolog 3
		Subclass: Receptor protein-tyrosine kinase erbB-3 (c-erbB3) (Tyrosine
		kinase-type cell surface receptor HER3) Subclass: v-erb-b2 erythroblastic leukemia viral oncogene homolog 2,
		ineuro/glioblastorna derived oncogene homolog; Avian erythroblastic
		leukemia viral (v-erb-b2) oncogene homolog 2; v-erb-b2 avian erythroblastic
		leukemia viral oncogene homolog 2 (neuro/glioblastoma derived oncogene
		homolog)
	 	Subclass: Similar to v-erb-b2 avian erythroblastic leukemia viral oncogene
		homolog 3
	İ	Subclass: herstatin
NM_021522	F:2.21 (5to19)	ubiquitin specific protease 14
NP 067497.1		
NM_007711		
_	E-2 2 (3/40 M)	chloride channel protein
INT U31/3/.1	F:2.2 (YtoM)	lemonde enginer protein

1	ı	Solution Children about a protein 2 (CIC 2)
		Subclass: Chloride channel protein 3 (ClC-3).
		Subclass: Chloride channel protein 4 (ClC-4). Subclass: chloride channel 5; Chloride channel-5
	 	Subclass: chloride channel 3; Chiolide channel-3
NTM 000245	F:2.2 (7to19)	hematopoietically expressed homeobox; proline-rich homeodomain-
NM_008245	F:2.2 (71019)	I
NP 032271.1		containing transcription factor
		Alternate: Similar to hematopoietically expressed homeobox
	<u> </u>	Alternate: me obox related protein
		1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1
AK003121	F:2.2 (5to19)	hypothetical protein MGC3279 similar to collectins
BAB22581.1		
		Alternate: collectin sub-family member 10; collectin liver 1; collectin 34
NM_016704	F:2.2 (5to19)	
NP 057913.1		complement component
	<u> </u>	Subclass: complement component C6
		Subclass: Complement component 6 precursor
		Subclass: similar to Complement component C6 precursor
		Subclass: complement C7
		Subclass: complement component 7 precursor
NM_021525	F:2.19 (7to19)	
NP 067500.1		RNA cyclase
112 00720012	<u> </u>	Subclass: RNA cyclase homolog
		Subclass: Similar to RNA cyclase homolog
		Subclass: RNA 3'-terminal phosphate cyclase-like protein
		Alternate: HSPC338
NM_022434	F:2.19 (5to19)	cytochrome P-450
NP 071879.1	ļ	
- 0,10,3,1		Subclass: cytochrome P450, subfamily IVF, polypeptide 2; leukotriene B4
		omega-hydrox ylase; leukotriene-B4 20-
		Subclass: cytochrome P450, subfamily IVA, polypeptide 11; fatty acid
	1	omega-hydrox ylase; P450HL-omega; alkane-1 monooxygenase; lauric acid
		omega-hydrox ylase
		Subclass: cytochrome P450, subfamily IVF, polypeptide 11
		Subclass: cytochrome P450 4F2
		Subclass: cytochrome P450, subfamily IVF, polypeptide 3; leukotriene B4
		lomega hydroxylase; leukotriene-B4 20-monooxygenase; cytochrome P450-
		LTB-omega
		Subclass: Cytochrome P450 4F12 (CYPIVF12)
		Subclass: similar to Cytochrome P450 4F12 (CYPIVF12) Subclass: cytochrome P450, subfamily IVF, polypeptide 8; microsomal
	<u> </u>	monooxygenase; flavoprotein-linked monooxygenase
		Subclass: similar to CYTOCHROME P450 4F6 (CYPIVF6)
		Alternate: hypothetical protein
		Alternate: F22329 1
		Alternate: Q9HBI6
NM_007899	F:2.18 (5to11)	extracellular matrix protein 1
NP 031925.1		
HAT 021372.1		1

			370
			Subclass: extracellular matrix protein 1, isoform 1 precursor; secretory
			component p85
			Subclass: extracellular matrix protein 1, isoform 2 precursor; secretory
			component p85
	NM_010028	F:2.18 (5to19)	
5	NP 034158.1	(3,015)	DEAD-box protein
5	NP U34136.1		Subclass: DEAD-box protein 3 (Helicase-like protein 2) (HLP2) (DEAD-
	·		box, X isoform)
			Subclass: DEAD/H (Asp-Glu-Ala-Asp/His) box polypeptide 3; DEAD/H
			box-3; heli case like protein 2; CAP-Rf
			Subclass: DEAD/H (Asp-Glu-Ala-Asp/His) box polypeptide 4; VASA
			protein
			Subclass: probable RNA helicase protein DKFZp434B1122.1
10			Subclass: similar to DEAD (aspartate-glutamate-alanine-aspartate) box
			polypeptide 3; D-E-A-D (aspartate-glutamate-alanine-aspartate) box
			polypeptide 3; embryonic RNA helicase
		·	Subclass: DEAD/H (Asp-Glu-Ala-Asp/His) box polypeptide 17 isoform 1;
			DEAD/H (Asp-Glu-Ala-Asp/His) box polypeptide 17 (72kD); probable
		·	RNA-dependent helicase p72
			Alternate: growth regulated nuclear 68 protein
	NM_053215	F:2.18 (5to19)	UDP glycosyltransferase; UDP-glucuronyltransferase
15	NP 444445.1		
	1,		Subclass: UDP glycosyltransferase 2 family, polypeptide B17; UDP-
	1	l ·	glucuronyl transferase, family 2, beta-17
•			Subclass: similar to UDP-glucuronosyltransferase 2B15 precursor,
			microsoma1 (UDPGT) (UDPGTH-3) (HLUG4)
		<u> </u>	Subclass: UDP glycosyltransferase 2 family, polypeptide B15; UDP-
		·	glucuronyl transferase, family 2, beta-15
			Subclass: UDP glycosyltransferase 2 family, polypeptide B4; UDP-
	ļ		glucuronyl transferase, family 2, beta-4 Subclass: similar to UDP-glucuronosyltransferase 2B4 precursor,
20			· · · · · · · · · · · · · · · · · · ·
		<u> </u>	microsoma1 (UDPGT) (Hyodeoxycholic acid) (HLUG25) (UDPGTH-1) Subclass: UDP glycosyltransferase 2 family, polypeptide B7; UDP-
	1.		
			glucuronyl transferase, family 2, beta-7 Subclass: UDP glycosyltransferase 2 family, polypeptide A1; UDP
,			glucuronos yltransferase 2 family, polypeptide A1
			Subclass: UDP glycosyltransferase 2 family, polypeptide B11
			Subclass: UDP glycosyltransferase 2 family, polypeptide B10
25			Subclass: UDP glycosyltransferase 2 family, polypeptide B28
	NM_015784		
•] —	T-0.17 (V:-0)	actocklast appecific factor 2 (faccialin I like); periostin
•	NP_056599.1	F:2.17 (Y to O)	osteoblast specific factor 2 (fasciclin I-like); periostin
30	AK007710	F:2.17 (5to19)	hypothetical protein FLJ12150
	BAB25204.1		
			Alternate: FKSG10

.

		371
NM 011415	F:2.16 (5to11)	snail 2; neural crest transcription factor SLUG; slug (chicken homolog),
NP 035545.1	` ´	finger protein
NP 035343.1		Alternate: snail 1 homolog; snail 1 zinc finger protein
		Alternate: similar to snail 1 (drosophila homolog), zinc finger protein
		Anteriale. Similar to said 1 (desception asserting)
AK011306	F:2.16 (5to19)	
BAB27532.1		eukaryotic translation initiation factor
		Subclass: eukaryotic translation initiation factor 5A; eIF5AI; eIF5A
		Subclass: similar to eukaryotic initiation factor 5A
		Subclass: eIF-5A2 protein; eIF5AII
NM 007686	F·2 16 (5to19)	Complement factor
	1.2.10 (3.01)	· ·
NP 031712.1	<u> </u>	
	<u> </u>	Subclass: Complement factor I Subclass: Complement factor I precursor (C3B/C4B inactivator)
	 	Subclass: Complement factor I precursor (C3B/C4B mactivalsi) Subclass: Similar to I factor (complement)
		outerass. Similar to Fractor (complement)
NM_010378	F:2.15	MHC class II antigen alpha chain
NP 034508.1	(11to19)	
		Subclass: MHC class II histocompatibility antigen HLA-DC-4 alpha cha
	,	precursor
		Subclass: MHC class II histocompatibility antigen HLA-DQ alpha 1 (DC
		specificity) precursor
		Subclass: HLA class II histocompatibility antigen, DQ(2) alpha chain
		Subclass: HLA class II histocompatibility antigen, DQ(5) alpha chain
		precursor (DC-1 alpha chain)
	 	Subclass: MHC HLA-DX-alpha chain
<u> </u>	 	HLA class II histocompatibility antigen, DQ(W3) alpha chain precursor
	 	Subclass: similar to HLA class II histocompatibility antigen, DQ(3) alph
		chain precursor (DC-alpha) (HLA-DCA) (HLA-DQA1*05011)
		Subclass: similar to HLA class II histocompatibility antigen, DP alpha cl
	1	
		precursor (HLA-SB alpha chain) (MHC class II DP3-alpha) (DP(W3))
	1	(DP(W4))
		Subclass: major histocompatibility complex, class II, DO alpha; lymphoc
	ŀ	antigen; HLA-D0-alpha; major histocompatibility complex, class II, DN
		alpha
····	 	Subclass: major histocompatibility complex, class II, DR alpha precursor
		HLA class II histocompatibility antigen, DR alpha chain
AK008273	F:2.15 (7to19)	Rho GDP dissociation inhibitor (GDI)
XP 132918.1		
		Subclass: Rho GDP dissociation inhibitor (GDI) beta; Ly-GDI
		Subclass: Rho GDP dissociation inhibitor (GDI) alpha
		Alternate: Ras-Related C3 Botulinum Toxin Substrate 2
ATZ010105	F-2 15 (5+010)	dynarnin 1-like protein
AK018195	F:2.13 (31019)	hynarimi 1-nke protein
BAC38054.1		Subclass: dynamin 1-like protein, isoform 1; dynamin-like protein

		372
		Sub class: dynamin 1-like protein, isoform 2; dynamin-like protein
		Subclass: dynamin 1-like protein, isoform 3; dynamin-like protein
		Sub class: Dnmlp/Vpslp-like protein
		Sub class: dynamin-like protein DYNIV-11
		Subclass: Similar to dynamin 1-like
		Subclass: dynamin, internal form 1, long C-terminal form
		Subclass: dynamin, internal form 2, short C-terminal form
		Subclass: Dynamin 2
		Sub class: dynamin 2; Dynamin II
		Subclass: similar to Dynamin 3 (Dynamin, testicular) (T-dynamin)
		Subclass: bA277C14.1 (novel Dynamin family member (KIAA0820))
		Alternate: KIAA0820 protein
NM_013562	F:2.15 (5to19)	
_	1	INTERFERON-RELATED DEVELOPMENTAL REGULATOR
NP 038590.1		Subclass: INTERFERON-RELATED DEVELOPMENTAL REGULATOR
	•	
	<u> </u>	1 (NERVE GROWTH FACTOR-INDUCIBLE PROTEIN PC4)
		Subclass: similar to INTERFERON-RELATED DEVELOPMENTAL
		REGULATOR 1 (NERVE GROWTH FACTOR-INDUCIBLE PROTEI)
	1	
,		PC4)
	 	Subclass: Interferon-related developmental regulator 2 (SKMC15 protein
NM 008015	F:2.14 (5to19)	DEAD box RNA helicase
	, ,	
NP_032041.1		Subclass: DEAD box RNA belicase DDX3
NP_032041.1		Subclass: DEAD box RNA helicase DDX3
NP_032041.1		Sub-class: dead box, X isoform
NP 032041.1		Subclass: dead box, X isoform Subclass: DEAD/H (Asp-Glu-Ala-Asp/His) box polypeptide, Y
NP_032041.1		Sub class: dead box, X isoform Sub class: DEAD/H (Asp-Glu-Ala-Asp/His) box polypeptide, Y chromosome; DEAD/H box-3, Y-linked
NP_032041.1		Subclass: dead box, X isoform Subclass: DEAD/H (Asp-Glu-Ala-Asp/His) box polypeptide, Y
NP_032041.1		Sub class: dead box, X isoform Sub class: DEAD/H (Asp-Glu-Ala-Asp/His) box polypeptide, Y chromosome; DEAD/H box-3, Y-linked
NP_032041.1		Subclass: dead box, X isoform Subclass: DEAD/H (Asp-Glu-Ala-Asp/His) box polypeptide, Y chromosome; DEAD/H box-3, Y-linked Subclass: similar to DEAD (aspartate-glutamate-alanine-aspartate) box polypeptide 3; D-E-A-D (aspartate-glutamate-alanine-aspartate) box
NP_032041.1		Subclass: dead box, X isoform Subclass: DEAD/H (Asp-Glu-Ala-Asp/His) box polypeptide, Y chromosome; DEAD/H box-3, Y-linked Subclass: similar to DEAD (aspartate-glutamate-alanine-aspartate) box polypeptide 3; D-E-A-D (aspartate-glutamate-alanine-aspartate) box polypeptide 3; embryonic RNA helicase
NP_032041.1		Subclass: dead box, X isoform Subclass: DEAD/H (Asp-Glu-Ala-Asp/His) box polypeptide, Y chromosome; DEAD/H box-3, Y-linked Subclass: similar to DEAD (aspartate-glutamate-alanine-aspartate) box polypeptide 3; D-E-A-D (aspartate-glutamate-alanine-aspartate) box polypeptide 3; embryonic RNA helicase Alternate: VASA protein
NP 052041.1		Subclass: dead box, X isoform Subclass: DEAD/H (Asp-Glu-Ala-Asp/His) box polypeptide, Y chromosome; DEAD/H box-3, Y-linked Subclass: similar to DEAD (aspartate-glutamate-alanine-aspartate) box polypeptide 3; D-E-A-D (aspartate-glutamate-alanine-aspartate) box polypeptide 3; embryonic RNA helicase
AK008590		Subclass: dead box, X isoform Subclass: DEAD/H (Asp-Glu-Ala-Asp/His) box polypeptide, Y chromosome; DEAD/H box-3, Y-linked Subclass: similar to DEAD (aspartate-glutamate-alanine-aspartate) box polypeptide 3; D-E-A-D (aspartate-glutamate-alanine-aspartate) box polypeptide 3; embryonic RNA helicase Alternate: VASA protein
AK008590	F:2.14 (5to19)	Subclass: dead box, X isoform Subclass: DEAD/H (Asp-Glu-Ala-Asp/His) box polypeptide, Y chromosome; DEAD/H box-3, Y-linked Subclass: similar to DEAD (aspartate-glutamate-alanine-aspartate) box polypeptide 3; D-E-A-D (aspartate-glutamate-alanine-aspartate) box polypeptide 3; embryonic RNA helicase Alternate: VASA protein Alternate: hypothetical protein
	F:2.14 (5to19)	Subclass: dead box, X isoform Subclass: DEAD/H (Asp-Glu-Ala-Asp/His) box polypeptide, Y chromosome; DEAD/H box-3, Y-linked Subclass: similar to DEAD (aspartate-glutamate-alanine-aspartate) box polypeptide 3; D-E-A-D (aspartate-glutamate-alanine-aspartate) box polypeptide 3; embryonic RNA helicase Alternate: VASA protein Alternate: hypothetical protein Ectonucleoside triphosphate diphosphohydrolase
AK008590	F:2.14 (5to19)	Subclass: dead box, X isoform Subclass: DEAD/H (Asp-Glu-Ala-Asp/His) box polypeptide, Y chromosome; DEAD/H box-3, Y-linked Subclass: similar to DEAD (aspartate-glutamate-alanine-aspartate) box polypeptide 3; D-E-A-D (aspartate-glutamate-alanine-aspartate) box polypeptide 3; embryonic RNA helicase Alternate: VASA protein Alternate: hypothetical protein Ectonucleoside triphosphate diphosphohydrolase Subclass: Ectonucleoside triphosphate diphosphohydrolase 1 (NTP Dase1
AK008590	F:2.14 (5to19)	Subclass: dead box, X isoform Subclass: DEAD/H (Asp-Glu-Ala-Asp/His) box polypeptide, Y chromosome; DEAD/H box-3, Y-linked Subclass: similar to DEAD (aspartate-glutamate-alanine-aspartate) box polypeptide 3; D-E-A-D (aspartate-glutamate-alanine-aspartate) box polypeptide 3; embryonic RNA helicase Alternate: VASA protein Alternate: hypothetical protein Ectonucleoside triphosphate diphosphohydrolase Subclass: Ectonucleoside triphosphate diphosphohydrolase 1 (NTP Dase1 (Ecto-ATP diphosphohydrolase) (ATPDase) (Lymphoid cell activation
AK008590	F:2.14 (5to19)	Subclass: dead box, X isoform Subclass: DEAD/H (Asp-Glu-Ala-Asp/His) box polypeptide, Y chromosome; DEAD/H box-3, Y-linked Subclass: similar to DEAD (aspartate-glutamate-alanine-aspartate) box polypeptide 3; D-E-A-D (aspartate-glutamate-alanine-aspartate) box polypeptide 3; embryonic RNA helicase Alternate: VASA protein Alternate: hypothetical protein Ectonucleoside triphosphate diphosphohydrolase Subclass: Ectonucleoside triphosphate diphosphohydrolase 1 (NTP Dase1 (Ecto-ATP diphosphohydrolase) (ATPDase) (Lymphoid cell activation antigen) (Ecto-apyrase) (CD39 antigen)
AK008590	F:2.14 (5to19)	Subclass: dead box, X isoform Subclass: DEAD/H (Asp-Glu-Ala-Asp/His) box polypeptide, Y chromosome; DEAD/H box-3, Y-linked Subclass: similar to DEAD (aspartate-glutamate-alanine-aspartate) box polypeptide 3; D-E-A-D (aspartate-glutamate-alanine-aspartate) box polypeptide 3; embryonic RNA helicase Alternate: VASA protein Alternate: hypothetical protein Ectonucleoside triphosphate diphosphohydrolase Subclass: Ectonucleoside triphosphate diphosphohydrolase 1 (NTP Dase1 (Ecto-ATP diphosphohydrolase) (ATPDase) (Lymphoid cell activation antigen) (Ecto-apyrase) (CD39 antigen) Subclass: Ectonucleoside triphosphate diphosphohydrolase 2 (NTP Dase2
AK008590	F:2.14 (5to19)	Subclass: dead box, X isoform Subclass: DEAD/H (Asp-Glu-Ala-Asp/His) box polypeptide, Y chromosome; DEAD/H box-3, Y-linked Subclass: similar to DEAD (aspartate-glutamate-alanine-aspartate) box polypeptide 3; D-E-A-D (aspartate-glutamate-alanine-aspartate) box polypeptide 3; embryonic RNA helicase Alternate: VASA protein Alternate: hypothetical protein Ectonucleoside triphosphate diphosphohydrolase Subclass: Ectonucleoside triphosphate diphosphohydrolase 1 (NTP Dase1 (Ecto-ATP diphosphohydrolase) (ATPDase) (Lymphoid cell activation antigen) (Ecto-apyrase) (CD39 antigen) Subclass: Ectonucleoside triphosphate diphosphohydrolase 2 (NTP Dase2 (Ecto-ATPase) (CD39 antigen-like 1)
AK008590	F:2.14 (5to19)	Subclass: dead box, X isoform Subclass: DEAD/H (Asp-Glu-Ala-Asp/His) box polypeptide, Y chromosome; DEAD/H box-3, Y-linked Subclass: similar to DEAD (aspartate-glutamate-alanine-aspartate) box polypeptide 3; D-E-A-D (aspartate-glutamate-alanine-aspartate) box polypeptide 3; embryonic RNA helicase Alternate: VASA protein Alternate: hypothetical protein Ectonucleoside triphosphate diphosphohydrolase Subclass: Ectonucleoside triphosphate diphosphohydrolase 1 (NTP Dase1 (Ecto-ATP diphosphohydrolase) (ATPDase) (Lymphoid cell activation antigen) (Ecto-apyrase) (CD39 antigen) Subclass: Ectonucleoside triphosphate diphosphohydrolase 2 (NTP Dase2 (Ecto-ATPase) (CD39 antigen-like 1) Subclass: ectonucleoside triphosphate diphosphohydrolase 3; CD39-like
AK008590	F:2.14 (5to19)	Subclass: dead box, X isoform Subclass: DEAD/H (Asp-Glu-Ala-Asp/His) box polypeptide, Y chromosome; DEAD/H box-3, Y-linked Subclass: similar to DEAD (aspartate-glutamate-alanine-aspartate) box polypeptide 3; D-E-A-D (aspartate-glutamate-alanine-aspartate) box polypeptide 3; embryonic RNA helicase Alternate: VASA protein Alternate: hypothetical protein Ectonucleoside triphosphate diphosphohydrolase Subclass: Ectonucleoside triphosphate diphosphohydrolase 1 (NTP Dase1 (Ecto-ATP diphosphohydrolase) (ATPDase) (Lymphoid cell activation antigen) (Ecto-apyrase) (CD39 antigen) Subclass: Ectonucleoside triphosphate diphosphohydrolase 2 (NTP Dase2 (Ecto-ATPase) (CD39 antigen-like 1)
AK008590 BAB25764.1	F:2.14 (5to19)	Subclass: dead box, X isoform Subclass: DEAD/H (Asp-Glu-Ala-Asp/His) box polypeptide, Y chromosome; DEAD/H box-3, Y-linked Subclass: similar to DEAD (aspartate-glutamate-alanine-aspartate) box polypeptide 3; D-E-A-D (aspartate-glutamate-alanine-aspartate) box polypeptide 3; embryonic RNA helicase Alternate: VASA protein Alternate: hypothetical protein Ectonucleoside triphosphate diphosphohydrolase Subclass: Ectonucleoside triphosphate diphosphohydrolase 1 (NTP Dase1 (Ecto-ATP diphosphohydrolase) (ATPDase) (Lymphoid cell activation antigen) (Ecto-apyrase) (CD39 antigen) Subclass: Ectonucleoside triphosphate diphosphohydrolase 2 (NTP Dase2 (Ecto-ATPase) (CD39 antigen-like 1) Subclass: ectonucleoside triphosphate diphosphohydrolase 3; CD39-like
AK008590 BAB25764.1	F:2.14 (5to19)	Subclass: dead box, X isoform Subclass: DEAD/H (Asp-Glu-Ala-Asp/His) box polypeptide, Y chromosome; DEAD/H box-3, Y-linked Subclass: similar to DEAD (aspartate-glutamate-alanine-aspartate) box polypeptide 3; D-E-A-D (aspartate-glutamate-alanine-aspartate) box polypeptide 3; embryonic RNA helicase Alternate: VASA protein Alternate: hypothetical protein Ectonucleoside triphosphate diphosphohydrolase Subclass: Ectonucleoside triphosphate diphosphohydrolase 1 (NTP Dase1 (Ecto-ATP diphosphohydrolase) (ATPDase) (Lymphoid cell activation antigen) (Ecto-apyrase) (CD39 antigen) Subclass: Ectonucleoside triphosphate diphosphohydrolase 2 (NTP Dase2 (Ecto-ATPase) (CD39 antigen-like 1) Subclass: ectonucleoside triphosphate diphosphohydrolase 3; CD39-like Subclass: E-type ATPase
AK008590 BAB25764.1	F:2.14 (5to19)	Subclass: dead box, X isoform Subclass: DEAD/H (Asp-Glu-Ala-Asp/His) box polypeptide, Y chromosome; DEAD/H box-3, Y-linked Subclass: similar to DEAD (aspartate-glutamate-alanine-aspartate) box polypeptide 3; D-E-A-D (aspartate-glutamate-alanine-aspartate) box polypeptide 3; embryonic RNA helicase Alternate: VASA protein Alternate: hypothetical protein Ectonucleoside triphosphate diphosphohydrolase Subclass: Ectonucleoside triphosphate diphosphohydrolase 1 (NTP Dase1 (Ecto-ATP diphosphohydrolase) (ATPDase) (Lymphoid cell activation antigen) (Ecto-apyrase) (CD39 antigen) Subclass: Ectonucleoside triphosphate diphosphohydrolase 2 (NTP Dase2 (Ecto-ATPase) (CD39 antigen-like 1) Subclass: ectonucleoside triphosphate diphosphohydrolase 3; CD39-like Subclass: E-type ATPase
AK008590 BAB25764.1	F:2.14 (5to19)	Subclass: dead box, X isoform Subclass: DEAD/H (Asp-Glu-Ala-Asp/His) box polypeptide, Y chromosome; DEAD/H box-3, Y-linked Subclass: similar to DEAD (aspartate-glutamate-alanine-aspartate) box polypeptide 3; D-E-A-D (aspartate-glutamate-alanine-aspartate) box polypeptide 3; embryonic RNA helicase Alternate: VASA protein Alternate: hypothetical protein Ectonucleoside triphosphate diphosphohydrolase Subclass: Ectonucleoside triphosphate diphosphohydrolase 1 (NTP Dase1 (Ecto-ATP diphosphohydrolase) (ATPDase) (Lymphoid cell activation antigen) (Ecto-apyrase) (CD39 antigen) Subclass: Ectonucleoside triphosphate diphosphohydrolase 2 (NTP Dase2 (Ecto-ATPase) (CD39 antigen-like 1) Subclass: ectonucleoside triphosphate diphosphohydrolase 3; CD39-like Subclass: E-type ATPase cytokine-inducible SH2-containing protein Subclass: cytokine-inducible SH2-containing protein isoform 2; cytokine
AK008590 BAB25764.1	F:2.14 (5to19)	Subclass: dead box, X isoform Subclass: DEAD/H (Asp-Glu-Ala-Asp/His) box polypeptide, Y chromosome; DEAD/H box-3, Y-linked Subclass: similar to DEAD (aspartate-glutamate-alanine-aspartate) box polypeptide 3; D-E-A-D (aspartate-glutamate-alanine-aspartate) box polypeptide 3; embryonic RNA helicase Alternate: VASA protein Alternate: hypothetical protein Ectonucleoside triphosphate diphosphohydrolase Subclass: Ectonucleoside triphosphate diphosphohydrolase 1 (NTP Dase1 (Ecto-ATP diphosphohydrolase) (ATPDase) (Lymphoid cell activation antigen) (Ecto-apyrase) (CD39 antigen) Subclass: Ectonucleoside triphosphate diphosphohydrolase 2 (NTP Dase2 (Ecto-ATPase) (CD39 antigen-like 1) Subclass: ectonucleoside triphosphate diphosphohydrolase 3; CD39-like . Subclass: E-type ATPase

			373
		1	Subclass: cytokine-inducible SH2-containing protein isoform 1; cytokine-
			inducible SH2-containing protein; cytokine-inducible inhibitor of signaling
		İ	type 1B; suppressor of cytokine signaling
l	1_025774	F:2.13 (5to11)	hypothetical protein FLJ13902
NP	080050.1		
NN	1 019775	F:2.13 (5to19)	
i	- 062749.1		plasma carboxypeptidase B
1	0027 13.1		Subclass: plasma carboxypeptidase B2 isoform a preproprotein; thrombin-
-			activatable fibrinolysis inhibitor; thrombin-activable fibrinolysis inhibitor;
	•		carboxypeptidase U; carboxypeptidase B-like protein; procarboxypeptidase
		ļ	U; procarboxypeptidase R; plasma procarboxypeptidase B
			Subclass: plasma carboxypeptidase B2 isoform b; thrombin-activable
			fibrimolysis inhibitor; thrombin-activatable fibrinolysis inhibitor;
			carboxypeptidase U; carboxypeptidase B-like protein; procarboxypep tidase
		·	U; procarboxypeptidase R; plasma procarboxypeptidase B
			Subclass: carboxypeptidase B-like protein
- 1	1_008348		
NP	032374.1	F:2.12 (YtoO)	inter1eukin 10 receptor, alpha
NM	1_020590	F:2.12 (7to19)	GABA(A) receptor-associated protein
NP	065615.1		
	-		Subclass: GABA(A) receptor-associated protein like 1; early estrogen-
			regulated protein
		:	Subclass: GABA(A) receptors associated protein like 3
NV.	1 011375	F:2.12 (5to19)	
1	035505.1		sialv l transferase
111	000000.1		Subclass: sialyltransferase 9 (CMP-NeuAc:lactosylceramide alpha-2, 3-
			sialyltransferase; GM3 synthase); ganglioside G(M3) Synthase
			Subclass: sialyltransferase 6 isoform j; Gal beta-1,3(4)GlcNAc alpha-2,3
			sialyltransferase; CMP-N-acetylneuraminate-beta-1,4-galactoside alpha-2,3-
			sialyltransferase; alpha-2,3-sialyltransferase II; alpha 2,3-sialyltransfer ase III
			in 1 2 and the control of the contro
- 1	1_013563		interleukin 2 receptor, gamma chain, precursor; Interleukin-2 receptor,
NP	038591.1	F:2.11 (YtoO)	gamma; common cytokine receptor gamma chain; CD132
NN	1_021291		
- [F:2.11 (YtoM)	amino acid transporter
-			Subclass: solute carrier family 7 (cationic amino acid transporter, y+
			system), member 9; solute carrier family 7, member 9; solute carrier family 7
	•		(cationic amino acid, transporter, y+ system), member 9
			Subclass: solute carrier family 7 (cationic amino acid transporter, y+
		1	system), member 5; Membrane protein E16; Solute carrier family 7, member

		374
<u> </u>		5; 4F2 light chain Subclass: solute carrier family 7, (cationic amino acid transporter, y+
ı		l ·
		system) member 11; cystine/glutamate transporter Subclass: solute carrier family 7 (cationic amino acid transporter, y+
<u> </u>		system), member 7 Subclass: solute carrier family 7 (cationic amino acid transporter, y+
 .		system), member 6 Subclass: solute carrier family 7 (cationic amino acid transporter, y+
	·	system), member 8
· · · · · · · · · · · · · · · · · · ·		Subclass: Y+L amino acid transporter 1 (y(+)L-type amino acid transporter
	1	1) (y+LAT-1) (Y+LAT1) (Monocyte amino acid permease 2) (MOP-2).
	<u> </u>	Subclass: solute carrier family 7, member 10; asc-type amino acid
		transporter 1
		Subclass: Large neutral amino acids transporter small suburnit 2 (L-type
		amino acid transporter 2) (hLAT2).
		arimio dold ranoporto: 2) (122-12-)
NM_010016	F:2.11 (7to11)	decay-acceleration factor
NP 034146.1		
		Subclass: decay accelerating factor for complement (CD55, Cromer blood
i i		group system); Decay-accelerating factor of
		Subclass: decay-accelerating factor, splice form 1 precursor
		Subclass: decay-accelerating factor 1 ab Subclass: decay-accelerating factor 4ab
		Subclass: decay-accelerating factor 3
L16846	F:2.11 (7to19)	B-cell translocation protein; Subclass: B-cell translocation protein 1
AAA37327.1		
	L .	IC 1 1 DTC formile manufaction 1 D coll temple control cone 1
i		Subclass: BTG family, member 2; B-cell translocation gene 2
		(pheochromacytoma cell-3); B-cell translocation gene 2
7, 000010	F. 11 (7, 10)	
	F:2.11 (7to19)	(pheochromacytoma cell-3); B-cell translocation gene 2
	F:2.11 (7to19)	(pheochromacytoma cell-3); B-cell translocation gene 2 Heat shock protein
	F:2.11 (7to19)	(pheochromacytoma cell-3); B-cell translocation gene 2 Heat shock protein Subclass: heat shock 70kDa protein 5 (glucose-regulated protein, 78kDa);
	F:2.11 (7to19)	(pheochromacytoma cell-3); B-cell translocation gene 2 Heat shock protein Subclass: heat shock 70kDa protein 5 (glucose-regulated protein, 78kDa); BiP; heat shock 70kD protein 5 (glucose-regulated protein, 78kD); Heat-
	F:2.11 (7to19)	(pheochromacytoma cell-3); B-cell translocation gene 2 Heat shock protein Subclass: heat shock 70kDa protein 5 (glucose-regulated protein, 78kDa); BiP; heat shock 70kD protein 5 (glucose-regulated protein, 78kD); Heat-shock 70kD protein-5 (glucose-regulated protein, 78kD)
	F:2.11 (7to19)	(pheochromacytoma cell-3); B-cell translocation gene 2 Heat shock protein Subclass: heat shock 70kDa protein 5 (glucose-regulated protein, 78kDa); BiP; heat shock 70kD protein 5 (glucose-regulated protein, 78kD); Heat-shock 70kD protein-5 (glucose-regulated protein, 78kD) Subclass: similar to 78 kDa glucose-regulated protein precursor (GRP 78)
	F:2.11 (7to19)	(pheochromacytoma cell-3); B-cell translocation gene 2 Heat shock protein Subclass: heat shock 70kDa protein 5 (glucose-regulated protein, 78kDa); BiP; heat shock 70kD protein 5 (glucose-regulated protein, 78kD); Heat-shock 70kD protein-5 (glucose-regulated protein, 78kD)
	F:2.11 (7to19)	(pheochromacytoma cell-3); B-cell translocation gene 2 Heat shock protein Subclass: heat shock 70kDa protein 5 (glucose-regulated protein, 78kDa); BiP; heat shock 70kD protein 5 (glucose-regulated protein, 78kD); Heat-shock 70kD protein-5 (glucose-regulated protein, 78kD) Subclass: similar to 78 kDa glucose-regulated protein precursor (GRP 78) (Immunoglobulin heavy chain binding protein) (BIP) (Endoplasmic reticulum lumenal Ca2+ binding protein grp78)
	F:2.11 (7to19)	(pheochromacytoma cell-3); B-cell translocation gene 2 Heat shock protein Subclass: heat shock 70kDa protein 5 (glucose-regulated protein, 78kDa); BiP; heat shock 70kD protein 5 (glucose-regulated protein, 78kD); Heat-shock 70kD protein-5 (glucose-regulated protein, 78kD) Subclass: similar to 78 kDa glucose-regulated protein precursor (GRP 78) (Immunoglobulin heavy chain binding protein) (BIP) (Endoplasmic reticulum lumenal Ca2+ binding protein grp78) Subclass: heat shock 70kDa protein 8 isoform 1; heat shock Cognate protein,
	F:2.11 (7to19)	Heat shock protein Subclass: heat shock 70kDa protein 5 (glucose-regulated protein, 78kDa); BiP; heat shock 70kD protein 5 (glucose-regulated protein, 78kD); Heat-shock 70kD protein-5 (glucose-regulated protein, 78kD) Subclass: similar to 78 kDa glucose-regulated protein precursor (GRP 78) (Immunoglobulin heavy chain binding protein) (BIP) (Endoplasmic reticulum lumenal Ca2+ binding protein grp78) Subclass: heat shock 70kDa protein 8 isoform 1; heat shock cognate protein, 71-kDa; heat shock 70kd protein 10; heat shock cognate protein 54;
	F:2.11 (7to19)	(pheochromacytoma cell-3); B-cell translocation gene 2 Heat shock protein Subclass: heat shock 70kDa protein 5 (glucose-regulated protein, 78kDa); BiP; heat shock 70kD protein 5 (glucose-regulated protein, 78kD); Heat-shock 70kD protein-5 (glucose-regulated protein, 78kD) Subclass: similar to 78 kDa glucose-regulated protein precursor (GRP 78) (Immunoglobulin heavy chain binding protein) (BIP) (Endoplasmic reticulum lumenal Ca2+ binding protein grp78) Subclass: heat shock 70kDa protein 8 isoform 1; heat shock Cognate protein,
	F:2.11 (7to19)	Heat shock protein Subclass: heat shock 70kDa protein 5 (glucose-regulated protein, 78kDa); BiP; heat shock 70kD protein 5 (glucose-regulated protein, 78kD); Heat-shock 70kD protein-5 (glucose-regulated protein, 78kD) Subclass: similar to 78 kDa glucose-regulated protein precursor (GRP 78) (Immunoglobulin heavy chain binding protein) (BIP) (Endoplasmic reticulum lumenal Ca2+ binding protein grp78) Subclass: heat shock 70kDa protein 8 isoform 1; heat shock cognate protein, 71-kDa; heat shock 70kd protein 10; heat shock cognate protein 54; constitutive heat shock protein 70; lipopolysaccharide-associated protein 1;
	F:2.11 (7to19)	Heat shock protein Subclass: heat shock 70kDa protein 5 (glucose-regulated protein, 78kDa); BiP; heat shock 70kD protein 5 (glucose-regulated protein, 78kD); Heat-shock 70kD protein-5 (glucose-regulated protein, 78kD) Subclass: similar to 78 kDa glucose-regulated protein precursor (GRP 78) (Immunoglobulin heavy chain binding protein) (BIP) (Endoplasmic reticulum lumenal Ca2+ binding protein grp78) Subclass: heat shock 70kDa protein 8 isoform 1; heat shock cognate protein, 71-kDa; heat shock 70kd protein 10; heat shock cognate protein 54; constitutive heat shock protein 70; lipopolysaccharide-associated protein 1; LPS-associated protein 1 Subclass: Heat shock 70 kDa protein 1 (HSP70.1) (HSP70-1/HSP70-2)
	F:2.11 (7to19)	Heat shock protein Subclass: heat shock 70kDa protein 5 (glucose-regulated protein, 78kDa); BiP; heat shock 70kD protein 5 (glucose-regulated protein, 78kD); Heat-shock 70kD protein-5 (glucose-regulated protein, 78kD) Subclass: similar to 78 kDa glucose-regulated protein precursor (GRP 78) (Immunoglobulin heavy chain binding protein) (BIP) (Endoplasmic reticulum lumenal Ca2+ binding protein grp78) Subclass: heat shock 70kDa protein 8 isoform 1; heat shock cognate protein, 71-kDa; heat shock 70kd protein 10; heat shock cognate protein 54; constitutive heat shock protein 70; lipopolysaccharide-associated protein 1;
NM_022310 NP_071705.1	F:2.11 (7to19)	Heat shock protein Subclass: heat shock 70kDa protein 5 (glucose-regulated protein, 78kDa); BiP; heat shock 70kD protein 5 (glucose-regulated protein, 78kD); Heat-shock 70kD protein-5 (glucose-regulated protein, 78kD) Subclass: similar to 78 kDa glucose-regulated protein precursor (GRP 78) (Immunoglobulin heavy chain binding protein) (BIP) (Endoplasmic reticulum lumenal Ca2+ binding protein grp78) Subclass: heat shock 70kDa protein 8 isoform 1; heat shock cognate protein, 71-kDa; heat shock 70kD protein 10; heat shock cognate protein 54; constitutive heat shock protein 70; lipopolysaccharide-associated protein 1; LPS-associated protein 1 Subclass: Heat shock 70kDa protein 1 (HSP70.1) (HSP70-1/HSP70-2) Subclass: heat shock 70kDa protein 1-like; Heat-shock 70kD protein-like-1; heat shock 70kD protein-like 1; heat shock 70kD protein 1-like
	F:2.11 (7to19)	Heat shock protein Subclass: heat shock 70kDa protein 5 (glucose-regulated protein, 78kDa); BiP; heat shock 70kD protein 5 (glucose-regulated protein, 78kD); Heat-shock 70kD protein-5 (glucose-regulated protein, 78kD) Subclass: similar to 78 kDa glucose-regulated protein precursor (GRP 78) (Immunoglobulin heavy chain binding protein) (BIP) (Endoplasmic reticulum lumenal Ca2+ binding protein grp78) Subclass: heat shock 70kDa protein 8 isoform 1; heat shock cognate protein, 71-kDa; heat shock 70kd protein 10; heat shock cognate protein 54; constitutive heat shock protein 70; lipopolysaccharide-associated protein 1; LPS-associated protein 1 Subclass: Heat shock 70 kDa protein 1 (HSP70.1) (HSP70-1/HSP70-2) Subclass: heat shock 70kDa protein 1-like; Heat-shock 70kD protein-like-1;

			375
		<u> </u>	Subclass: heat shock 70kDa protein 1B; heat shock 70kD protein 1B
			Subclass: heat shock 70kDa protein 2; heat shock 70kD protein 2; Heat-
			shock 70kD protein-2 Subclass: heat shock cognate protein 54
			Subclass: heat shock cognate protein 94 Subclass: heat shock 70kDa protein 9B (mortalin-2); heat shock 70kD
			protein 9 (mortalin); mot-2; mthsp75; heat shock 70kD protein 9B (mortalin-
			2); Heat-shock 70kD protein-9 (mortalin)
5			Subclass: heat shock 70kDa protein 6 (HSP70B'); heat shock 70kD protein 6
			(HSP70B'); Heat-shock 70kD protein-6 (HSP70B')
		 	Subclass: Similar to heat shock cognate 71-kd protein
			Subclass: dnaK-type molecular chaperone HSPA1L Alternate: Unknown (protein for MGC:33922)
			·
10	AK004654	F:2.11 (5to7)	Similar to hypothetical protein FLJ13511
	BAB23445.1		
			Alternate: F02569 2 Alternate: 7h3 protein
		<u> </u>	Attendate. 7115 protein
15	AK009563	F:2.1 (5to19)	Protein KIAA1434
	BAB26361.1		
			Alternate: similar to KIAA1434 protein
			Alternate: unnamed protein product
20	NM_011579	F:2.1 (5to19)	hypothetical protein R30953 1
	NP 035709.1		
	141 055705.1		
	NM_021394	F:2.1 (5to19)	dJ718J7.3.1 (novel protein similar to mouse tumour stroma and activated
	NP_067369.1		macrophage protein DLM-1, isoform 1)
25.			Alternate: tumor stroma and activated macrophage protein DLM-1;
			chromosome 20 open reading frame 183
	NM 016702	F:2.1 (5to19)	alanine-glyoxylate aminotransferase; alanine-glyoxylate aminotransferase,
	, –	1.2.1 (51019)	
	NP_057911.1		liver-specific peroxisomal; serine-pyruvate aminotransferase
30 ຶ	NM_013550		
	NP_038578.1	F:2.09 (YtoO)	H4 histone family, member A
	AK003938		
~ ~	BAB23084.1	F:2.09 (YtoM)	KIAA1866 protein
35		 	
			tetraspan 5; tetraspan TM4SF; tetraspan NET-4; transmembrane 4
	NM_019571		superfamily member 9; transmembrane 4 superfamily, member 8; tetraspanin
	1 -	E-2 00 (V+2) (1)	
	INF 002317.1	F:2.09 (YtoM)	2
40	NM_007509	F:2.09 (7to19)	
	NP 031535.2		ATPase
			Subclass: ATPase, H+ transporting, lysosomal 56/58kD, V1 subunit B,

		376
1		58 kDa subunit; vacuolar ATP synthase subunit B, brain is oform; V-ATPase
		B2 subunit; H(+)-transporting two-sector ATPase, 56/58kD subunit, isoform
	<u> </u>	2
		Subclass: ATPase, H+ transporting, lysosomal 56/58kD, V1 subunit B,
		isoform 1; ATPase, H+ transporting, lysosomal, beta polypeptide, 58kD;
	1	vacuolar proton pump, subunit 3; vacuolar ATP synthase subunit B, kidney
		isoform; V-ATPase B1 subunit; endomembrane proton purnp 58 kDa
		subunit; H(+)-transporting two-sector ATPase, 58kD suburnit; H+-ATPase
		beta 1 subunit; ATPase, H+ transporting, lysosomal 56/58kcD, V1 subunit B,
		isoform 1 (Renal tubular acidosis with deafness)
		Alternate: Unknown (protein for IMAGE:3352651)
		Alternate: Unknown (protein for MGC:32642)
ND (000022	E-2 00 (7to10)	prolactin receptor
NM_008932	F:2.09 (7to19)	protactini receptor
NP_032958.1		Cub along, muslo atim magantan
		Subclass: prolactin receptor Subclass: prolactin receptor short isoform 1a
		Subclass: intermediate prolactin receptor isoform
		Subclass: prolactin receptor short isoform 1b
		Subclass: delta 4-SF1b truncated prolactin receptor
		Subclass: prolactin receptor isoform delta S1 precursor
		Subclass: delta 4-delta 7/11 truncated prolactin receptor
		Subciass, deta 4-deta //11 transacta pronociii 1000ptoi
AK003950	F:2.09 (5to19)	Similar to RIKEN cDNA 1110029A09 gene
BAB23088.1	, ,	
BAB23088.1		Alternate: unnamed protein product
AK010325	F:2.09 (5to19)	
NP_542123.1		transmembrane protein 9 superfamily
		Subclass: transmembrane 9 superfamily member 1; multispanning membrane
		protein (70kD); transmembrane protein 9 superfamily member 1
		Subclass: transmembrane 9 superfamily member 2; 76 kDa membrane
		protein; transmembrane protein 9 superfamily member 2
		Subclass: transmembrane protein TM9SF3
		Subclass: similar to Transmembrane 9 superfamily protein member 3
		precursor (SM-11044 binding protein) (EP70-P-iso)
	 	Alternate: KIAA0255 gene product
		Alternate: SM-11044 binding protein
		Alternate: unnamed protein product
		·
NM_011521	F:2.09 (5to19)	Syndecan-4 precursor (Amphiglycan) (SYND4) (Ryudocara core protein)
NP_035651.1		
NTM 010427	E-2 00 (5+c10)	hypothetical protein FLJ11149
NM_019437	(21019)	hisponionon protein i Esti 142
NP 062310.1		A1,
		Alternate: unnamed protein product
NM 007811	F:2.08 (5to11)	Cytochrome P450 26 (Retinoic acid-metabolizing cytochrome) (P450RAI)
		l i
NP_031837.1	L	(hP450RAI) (Retinoic acid 4-hydroxylase)

-		. 377
	2000(5: 11)	
NM_010324	F:2.08 (5to1 1)	transaminase
NP_034454.1		
		Subclass: aspartate aminotransferase 1; glutamic-oxaloacetic transaminase 1
1		soluble
		Subclass: glutamic-oxaloacetic transaminase 2, mitochondrial (aspartate
		aminotransferase 2)
NM_008364	F:2.08 (5to19)	
NP 032390.1		interleukin 1 receptor accessory protein
		Subclass: interleukin 1 receptor accessory protein isoform 1
		Subclass: interleukin 1 receptor accessory protein isoform 2
•		Subclass: interleukin 1 receptor accessory protein-like 2; interleukin 1
		receptor 9; IL-1 receptor; X-linked interleukin-1 receptor accessory protein-
	İ	like 2; IL-1 receptor accessory protein-like 2
NM_023580	F:2.08 (5to 19)	receptor protein-tyrosine kinase
NP 076069.1		
		Subclass: EphA1; eph tyrosine kinase 1 (erythropoietin-producing hepatoma
		amplified sequence; oncogene EPH; ephrin receptor EphA1); eph tyrosine
		kinase 1 (erythropoietin-producing hepatoma amplified sequence); ephrin
		receptor EphA1
		Subclass: EphA2; ephrin receptor EphA2; epithelial cel1 receptor protein
		tyrosine kinase
		Subclass: EphA7; Hek11; ephrin receptor EphA7 Subclass: EphA4; Hek8; TYRO1 protein tyrosine kinase; ephrin receptor
İ		
		EphA4
		Subclass: Ephrin type-A receptor 5 precursor (Tyrosine-protein kinase
		receptor EHK-1) (Eph homology kinase-1) (Receptor protein-tyrosine kinase
		HEK7)
		Subclass: similar to Ephrin type-A receptor 5 precursor (Tyrosine-protein
		kinase receptor EHK-1) (Eph homology kinase-1) (Receptor protein-tyrosine
		kinase HEK7)
-		Subclass: EphA3; Ephrin receptor EphA3 (human embryo kinase 1); eph-like
,	1	tyrosine kinase 1 (human embryo kinase 1); ephrin receptor EphA3
 		Subclass: ephrin receptor EphA8 precursor; ephrin type-A receptor 8
	1	precursor; eph- and elk-related tyrosine kinase; tyrosylprotein kinase;
		<u> </u>
<u> </u>		tyrosine-protein kinase receptor eek; protein-tyrosine kimase; hydroxyaryl-
		protein kinase
		Subclass: ephrin receptor EphB6 precursor; tyrosine-protein kinase-defective
		receptor; ephrin type-B receptor 6
		Subclass: ephrin receptor EPHA3 secreted form
		Subclass: large erk kinase
		Subclass: dJ74M1.1.1 (tyrosine kinase isoform 1)
	<u> </u>	Subclass: dJ74M1.1.2 (tyrosine kinase isosform 2)
	 	Alternate: KIAA1459 protein
	<u> </u>	1

		378
AK004731		
XP 148015	F:2.07 (YtoM)	nlakophilin
AF 148015	1.2.07 (1 101/1)	Subclass: plakophilin 2
	 	Subclass: plakophilin 2a
- 		Subolass. Planoplatal 24
NM 008961	F-2.07 (7to19)	phosphotriesterase related; resiniferatoxin-binding, phosphotriesterase-
	[, (,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,	f -
NP 032987.1		related gene; phosphotriesterase-related
ND 4 025449	E-2.07 (7+=10)	signal sequence receptor, beta (translocon-associated protein beta)
NM_025448	F:2.07 (71019)	Signal Sequence receptor, beta (transfocon-associated protein beta)
NP_079724.1		
7.07.430	T.0.07 (51.10)	C (alaba) Protein C alaba
L27439	F:2.07 (5to19)	protein S (alpha); Protein S, alpha
AAA40006.1		
		Alternate: growth arrest-specific 6; AXL stimulatory factor
	5005(5: 10)	1 1 701 D
M12573	F:2.07 (5to19)	heat shock 70kDa protein
AAA37863.1		
		Subclass: heat shock 70kDa protein 1A; heat shock 70kD protein 1A; heat
		shock-induced protein; dnaK-type molecular chaperone HSP70-1
		Subclass: heat shock 70kDa protein 1B; heat shock 70kD protein 1B
		Subclass: heat shock 70kD protein 1-like
		Subclass: heat shock 70kDa protein 6 (HSP70B')
		Subclass: heat shock 70kDa protein 2; heat shock 70kD protein 2; Heat-
	[shock 70kD protein-2
·_		Subclass: heat shock 70kDa protein 8 isoform 1; heat shock cognate protein
		1
		71-kDa; heat shock 70kd protein 10; heat shock cognate protein 54;
		constitutive heat shock protein 70; lipopolysaccharide-associated protein 1;
• •		LPS-associated protein 1
		Subclass: Similar to heat shock 70kD protein 8
		Subclass: similar to HEAT SHOCK 70 KD PROTEIN 6 (HEAT SHOCK
•	· ·	KD PROTEIN B)
		RD FROTEIN B)
NM 009780	F-2 07 (5to19)	complement component
_	1.2.07 (3.013)	bompomon component
NP_033910.1	·	
		Subclass: complement component C4
		Subclass: complement component C4A Subclass: complement component 4A preproprotein; acidic C4; Rodgers
	1	1
		form of C4; complement component 4S
		Subclass: complement component 4B preproprotein; Chido form of C4;
		basic C4; complement component 4F
		Subclass: complement component C4B
		Subclass: complement C4d
		Subclass: complement C4d variant
		Subclass: complement component 3
		Subclass: complement component 5
U27315	F:2.07 (5to19)	
AAC52837.1	1	solute carrier family (mitochondrial carrier; adenine nucleotide translocate
	1	pointe carrier rainity (introducidatal carrier, adomite incidende translocate

. 75

		379
		translocator), member 4; adenine nucleotide translocator 1
		Subclass: similar to ADP,ATP carrier protein, heart/skeletal muscle isofor
		T1 (ADP/ATP translocase 1) (Adenine nucleotide translocator 1) (ANT1)
		Subclass: solute carrier family 25 (mitochondrial carrier; adenine nucleot
		translocator), member 5; 2F1; adenine nucleotide translocator 2
		Subclass: ADP/ATP carrier protein (adenine nucleotide translocator 2)
		Subclass: similar to ADP, ATP carrier protein, fibroblast isoform (ADP/A
		translocase 2) (Adenine nucleotide translocator 2) (ANT 2)
		Subclass: similar to ADP,ATP carrier protein, liver isoform T2 (ADP/AT
		translocase 3) (Adenine nucleotide translocator 3) (ANT 3)
		Subclass: Similar to solute carrier family 25 (mitochondrial carrier; adeni
		nucleotide translocator), member 5
		Alternate: hypothetical protein DKFZp434N1235
NM_007860	F:2.O6 (7to19)	Type I iodothyronine deiodinase (Type-I 5'deiodinase) (DIOI) (Type 1 DI
NP 031886.1		(5DI)
		Alternate: Similar to deiodinase, iodothyronine, type I
NM_020001	F:2.O6 (5to19)	dendritic cell lectin b; blood dendritic cell antigen 2 protein
- .	1.2.00 (3.0.1)	
NP_064385.1		
NM_026533	F:2.O6 (5to19)	ribosomal protein S13; 40S ribosomal protein S13
NP 080809.1		
NM_033373		
NP 203537.1	F:2.O5 (YtoO)	keratin
		Subclass: keratin 23 isoform a; hyperacetylation-iriducible type I keratin;
		keratin, type I cytoskeletal 23; cytokeratin 23; type I intermediate filamen
		cytokeratin; histone deacetylase inducible keratin 23, (Cytokeratin 23) (K
	}	(CK 23).
		Subclass: keratin 23 isoform b; hyperacetylation-inducible type I keratin;
		keratin, type I cytoskeletal 23; cytokeratin 23; type I intermediate filamen
,		cytokeratin; histone deacetylase inducible keratin 23
		Subclass: keratin 20, type I-like, cytoskeletal
		Subclass: keratin 19; keratin, type I cytoskeletal 19; keratin, type I, 40-kd
		cytokeratin 19; (Cytokeratin 19) (K19) (CK 19).
		Subclass: keratin 17
		Subclass: keratin 12 (Meesmann corneal dystrophy); Keratin-12; keratin 1
		Subclass: keratin 15; keratin-15, basic; keratin-15, beta; type I cytoskeleta
		15; cytokeratin 15; (Cytokeratin 15) (K15) (CK 15).
		Subclass: keratin 13; keratin, type I cytoskeletal 13; cytokeratin 13
		Subclass: keratin 16; keratin, type I cytoskeletal 16; cytokeratin 16
	<u> </u>	Subclass: keratin 14; cytokeratin 14
		Subclass: type I hair keratin 6; keratin, hair, acidic, 6
	 	Subclass: cytokeratin 20
		Subclass: type I hair keratin 5; Ha-5; hard keratin, type I, 5
	 	Subclass: Keratin, type I cytoskeletal 10 (Cytokeratin 10) (K10) (CK 10). Subclass: type I hair keratin 3A; Ha-3I; hard keratin, type I,3I; keratin, ha
-		acidic,3A
1		land and a second

•		380
	ł	Subclass: type I hair keratin 1; hard keratin, type I, 1; Ha-1; keratin, hair,
		acidic,1
		Subclass: type I hair keratin 4; hard keratin, type I, 4
		Subclass: type I hair keratin 2; Ha-2; hard keratin, type I, 2; keratin, hair,
		acidic,2
		Subclass: keratin 18
ļ		Subclass: cytokeratin 9
AK009020		
AK009020		·
BAB26030.2	F:2.05 (YtoM)	chloride intracellular channel
		Subclass: chloride intracellular channel 3
		Subclass: Chloride intracellular channel protein 2 (XAP121).
	_	Subclass: p64 bovine chloride channel-like protein
	<u> </u>	Subclass: chloride intracellular channel 1; p64CLCP
		Subclass: Chloride intracellular channel protein 5
		Subclass: chloride intracellular channel 4
		Subclass: chloride intracellular channel 6; chloride channel form A
		Subclass: H1 chloride channel; p64H1; CLIC4
<u></u>		Subclass: chloride channel form B
NM_025939	F-2 05 (7to 11)	phosphoribosylaminoimidazole carboxylase, phosphoribo sylaminoimidazole
_	1,2.03 (7011)	
NP_080215.1	1 .	succinocarboxamide synthetase; phosphoribosylaminoimi dazole carboxylase
	<u> </u>	phosphoribosylaminoribosylaminoimidazole succinocarboxamide synthetase
		Alternate: similar to Multifunctional protein ADE2
70.460.4	7 2 2 2 2 2 2 2 2 2 2 2 2 2 2 2 2 2 2 2	
J04694	F:2.05 (5to11)	collagen
AAA50292.1		
		Subclass: alpha 1 type IV collagen preproprotein; collagen IV, alpha-1
		polypeptide; collagen of basement membrane, alpha-1 chain
		Subclass: alpha-2 type IV collagen
		Subclass: alpha-3 type IV collagen
		Subclass: alpha 3 type IV collagen, isoform 1, precursor; collagen IV, alpha-
	1	3 polypeptide (goodpasture antigen)
ļ	 	Subclass: alpha-5 type IV collagen
	<u> </u>	Subclass: alpha 5 type IV collagen, isoform 2, precursor; collagen IV, alpha-
	1	
	 	5 polypeptide; collagen of basement membrane, alpha-5 chain
		Subclass: alpha 5 type IV collagen, isoform 3, precursor; collagen IV, alpha-
	<u> </u>	5 polypeptide; collagen of basement membrane, alpha-5 chain
		Subclass: alpha 5 type IV collagen, isoform 1, precursor; collagen IV, alpha-
		5 polypeptide; collagen of basement membrane, alpha-5 chain
		Subclass: type IV alpha 6 collagen, isoform A precursor; collagen IV, alpha-
		6 polypeptide; collagen of basement membrane, alpha-6
	1-	Alternate: arresten
		Alternate: tumstatin
	 	A ALOCA COMPANY
U70139	F:2.05 (5to7)	nocturnin
1	1.2.55 (5.67)	,
AAB62717.1		All CODA 1 and the literature in A liter (Const.) CODA
		Alternate: CCR4 carbon catabolite repression 4-like (S. cerevisiae); CCR4-
		like (carbon catabolite repression 4, S. cerevisiae)
1		

		381
MM_008956	IF :2.05 (5to19)	
NP 032982.1		polypyrimidine tract binding protein
·		Subclass: polypyrimidine tract binding protein, isoform c; RNA binding
		protein; heterogeneous nuclear ribonucleoprotein polypeptide I;
		polypyrimidine tract binding protein (heterogeneous nuclear
		ribonucleoprotein I)
	<u> </u>	Subclass: polypyrimidine tract binding protein, isoform b; RNA binding
		protein; heterogeneous nuclear ribonucleoprotein polypeptide I;
		polypyrimidine tract binding protein (heterogeneous nuclear
		ribonucleoprotein I)
		Subclass: polypyrimidine tract binding protein, isoform a; RNA binding
		protein; heterogeneous nuclear ribonucleoprotein polypeptide I;
		polypyrimidine tract binding protein (heterogeneous nuclear
		ribonucleoprotein I)
		Subclass: polypyrimidine tract binding protein 2; neural polypyrimidine trac
		binding protein; PTB-like protein
		Subclass: imilar to polypyrimidine-tract binding protein
		Subclass: non-neuronal splice variant nPTB 3
		Subclass: non-neuronal splice variant nPTB4
	_	Subclass: PTB-like protein L
		Subclass: PTB-like protein S
		1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1
NM_011919	F :2.05 (5to19)	inhibitor of growth family, member 1; inhibitor of growth 1; inhibitor of
NP 036049.1	.	growth 1 family, member 1
		Alternate: candidate tumor suppressor p33ING1
		Alternate: p47
		Alternate: p33
		Alternate: p24 is an alternatively spliced transcript of p33/ING1.
		Alternate: p32 protein
7 . 010 1/7	7 2 05 (5) 10)	I C. A. (IICE) activator
NM_019447	1F:2.05 (5to19)	hepatocyte growth factor (HGF) activator
NP 062320.1		
0.57206	T-2 05 (5+-10)	serum deprivation response protein; serum deprivation response;
S67386	F:2.05 (51019)	1
AAB28953.1		phosphatidylserine-binding protein
		Alternate: leucine-zipper protein FKSG13
NM_008039		
NP 032065.1	F:2.04 (YtoO)	N-formyl peptide receptor Subclass: formyl peptide receptor-like 1; lipoxin A4 receptor (formyl peptide)
		receptor related) Subclass: FMLP-related receptor II
<u> </u>		Subclass: formyl peptide receptor-like 2
ļ		
		fSubclass: ormyl peptide receptor 1
		Subclass: fMet-Leu-Phe receptor (fMLP receptor) (N-formyl peptide
		receptor) (FPR) (N-formylpeptide chemoattractant receptor).
		Subclass: N-formylpeptide receptor fMLP-R_98
L		

		382
VM_009417		
TD 0334/43 T	F:2.04 (YtoM)	Peroxidase
YE U33443.1	1.2.04 (1.0.1/1)	Subclass: thyroid peroxidase isoform a; thyro peroxidase; thyroid microso
		antigen
		Subclass: thyroid peroxidase isoform b, thyroperoxidase; thyroid microso
		antigen Subclass: thyroid peroxidase isoform c; thyro peroxidase; thyroid microsc
		antigen
		Subclass: thyroid peroxidase isoform d; thyro peroxidase; thyroid microsc
	•	antigen
		Subclass: thyroid peroxidase isoform e; thyroperoxidase; thyroid microso
		antigen
		Subclass: thyroid peroxidase isoform 5
		Subclass: myeloperoxidase
		Subclass: eosinophil peroxidase
		Subclass: lactoperoxidase
VM_007472	F:2.04 (7to11)	aquaporin (water channel protein)
NP 031498.1		·
T. ODITIONAL		Subclass: aquaporin 1 (channel-forming integral protein, 28kD)
		Subclass: major intrinsic protein of lens fiber; aquaporin
·		Subclass: aquaporin 2; Aquaporin-2 (collecting duct)
		Subclass: hAQP-CD=collecting duct aquaporin [human, kidney, Peptide
		271 aa]
		Subclass: aquaporin 4 C2 isoform; mercurial-insensitive water channel
	"	Subclass: aquaporin 4 isoform a; mercurial-imsensitive water channel
		Subclass: aquaporin 4, long splice form - hurman
		Subclass: aquaporin 5; Aquaporin-5
NM 029239	F:2.04 (7to11)	
	, í	protein kinase
NP 083515.1		Subclass: protein kinase C, nu; serine-threonime protein kinase
	ļ	Subclass: Similar to protein kinase C, nu
	ļ — — — — — — — — — — — — — — — — — — —	
		Subclass: protein kinase C, mu
	ļ	Subclass: protein kinase D2
AK003830	F:2.04 (7to19)	CGI-128 protein
BAB23024.1		
NB 4 000500	E-2 04 (7+a10)	carnitine/acylcarnitine translocase; Carnitine-acylcarnitine translocase;
NM_020520	F:2.04 (71019)	<u> </u>
NP_065266.1	1	carnitine-acylcarnitine carrier; solute carrier family 25
		(carnitine/acylcarnitine translocase), member 20
	F-2 04 (5to19)	uridine phosphorylase
えなののファム 4	1.2.04 (31013)	
AK007264		1
AK007264 BAB24924.1		
		Alternate: similar to Uridine phosphorylase (UDRPase)
BAB24924.1	F:2.04 (5to19)	Alternate: similar to Uridine phosphorylase (UDRPase) seven transmembrane domain protein
	F:2.04 (5to19)	

thinemia-hyperammonemia-	_011017 F:2.04 (5to19) 035147.1
thinemia-hyperammonemia-	035147.1
thinemia-hyperammonemia-	
	· ·
	_029796 F:2.04 (5to19)
	- ` `
	084072.1
	_021532
ene 3	1
she 5	067507.2 F:2.03 (YtoO)
	_011087
	_
ceptor, subfamily B (with TM	035217.1 F:2.03 (YtoO)
nmunoglobulin-like receptor	
variant 3	
variant 4	
ceptor-2	
ceptor, subfamily B (with TM	
nmunoglobulin-like receptor	
ceptor, subfamily A (without	
ulin-like receptor 4	·
ceptor, subfamily B (with TM	
ceptor, subfamily A (with TM	
ulin-like receptor 6	
ceptor, subfamily A (with TM	
ulin-like receptor 7	
nrotein	
J. C. CO. L.	
	019922 F:2.03
	- '
gilar to rat synaptonemal	064306.1 (11t019)
mai to rat by supro	-
	205602 10:202 (7:-11)
	580082 F:2.03 (7011)
	K62363.1
1-1 1	
in-like, hormone receptor-like	1 .
	ľ
in-like, hormone receptor-like	
n-like, hormone receptor-like	
protein nilar to rat synaptonemal	[_019922 F:2.03 064306.1 (11to19) 385682 F:2.03 (7to11) K62363.1

		384
1		Subclass: egf-like module containing, mucin-like, hormone receptor-like
		sequence 2 isoform b
	<u> </u>	Subclass: egf-like module containing, mucin-like, hormone receptor-like
ĺ	Ĭ	
	 	sequence 2 isoform c Subclass: egf-like module containing, mucin-like, hormone receptor-like
		1
		sequence 2 isoform e
İ		Subclass: egf-like module containing, mucin-like, hormone receptor-like
		sequence 2 isoform d
		Subclass: egf-like module containing, mucin-like, hormone receptor-like
		cogyomas 2 isoform a
		sequence 2 isoform g Subclass: egf-like module containing, mucin-like, hormone receptor-like
		}
		sequence 2 isoform f
	ļ	Subclass: egf-like module-containing mucim-like receptor 3 isoform a
		Subclass: EGF-like module EMR2
	<u> </u>	Alternate: lectomedin
	 	Subclass: lectomedin-3 Subclass: latrophilin 1; KIAA0786 protein; lectomedin-1; latrophilin
	1	Subclass: ladopmin 1; KIAAU/80 protein; rectornedii-1; radopmini Subclass: lectomedin-1 alpha
	 	vlectomedin-2
	<u> </u>	Subclass: lectomedin-2; KIAA0821 protein
		Alternate: CD97 antigen, isoform 1 precurs or; leukocyte antigen CD97;
		•
	 	seven-span transmembrane protein Alternate: CD97 antigen, isoform 2 precurs or; leukocyte antigen CD97;
		Alternate: CD97 antigen, isoform 2 precuis of; feukocyte antigen CD97,
		seven-span transmembrane protein
NM 008625	F:2.03 (7to19)	Mannose receptor C; Subclass: mannose receptor C type 1; mannose
-	(,	1
NP_032651.1	 	receptor precursor; macrophage mannose receptor Subclass: mannose receptor, C type 2; KIA A0709 gene product; endocytic
		·
		receptor (macrophage mannose receptor famuly); likely ortholog of mouse
		mannose receptor, C type 2
		Alternate: endocytic receptor Endo 180
		Alternate: phospholipase A2 receptor 1, 18OkDa; phospholipase A2 receptor
		1, 180kD
	 	1, 160KD
NM_008991	F:2.03 (7to19)	
_	(1102)	16 7 5
NIP 033017.1	ļ	ATP-binding cassette, sub-family D
		Subclass: ATP-binding cassette, sub-family D, member 3; Peroxisomal
		membrane protein-1 (70kD); peroxisomal membrane protein 1 (70kD,
	ļ	Zellweger syndrome); peroxisomal membrarne protein-1
		Subclass: ATP-binding cassette, sub-family D (ALD), member 1;
	1	1
		adrenoleukodystrophy protein
		Subclass: ATP-binding cassette, sub-family D, member 2;
		adrenoleukodystrophy-like 1; hALDR
NTN 4 025422	F.2 02 (7to 10)	KIA A 0022 gaza product
NIM_025422	r:2.03 (7019)	KIAA0022 gene product
NP 079698.1		
	<u> </u>	
NIM_007624	F:2.03 (5to19)	chromobox homolog 3; heterochromatin protein HP1 gamma; HP1 gamma

•	•	385
NP_031650.1		homolog; heterochromatin-like protein 1
NM 013762	F:2.03 (5to19)	
_	F.2.03 (31019)	i
NP_038790.1		ribosomal protein
		Subclass: ribosomal protein L3; 60S ribosomal protein L3; HIV-1 TAR
		RNA-binding protein B
		Subclass: similar to ribosomal protein L3; 60S ribosomal protein L3; H
-	i	TAR RNA-binding protein B
		Alternate: Unknown (protein for IMAGE:3538792)
		Alternate: ARBP-b gene product
		Alternate: Similar to RIKEN cDNA 1110057H16 gene
NTM 012927	E-2 02 (5+a10)	tyrosylprotein sulfotransferase; Subclass: tyrosylprotein sulfotransferas
NM_013837	F.2.03 (31019)	lytosymptotem sumon amsterase, is crociass. Tyrosymptotem sumon amsteras
NP_038865.1		
		Subclass: tyrosylprotein sulfotransferase 2; Tyrosylprotein
		phosphotransferase 2
		Alternate: hypothetical protein
ND 6 01 6751	E-2 02 (5) 10)	
NM_016751	F:2.03 (5to19)	similar to Kupffer cell receptor
NP_058031.1		
		Alternate: unnamed protein product
		Alternate: Langerhans cell specific c-type lectin; langerin
NTM 000042		
NM_008043		
NP_032069.1	F:2.02 (YtoM)	frequently rearranged in advanced T-cell lymphomas; FRAT1
NM 008905	F·2 02 (7to19)	hypothetical protein
-	1.2.02 (7.01)	hypothetical protein
NP_032931.1		Alternate: protein tyrosine phosphatase, receptor-type, F interacting pro
•		
		binding protein 2
	ļ	Alternate: similar to hypothetical protein
		Alternate: liprin-beta2 Alternate: PTPRF interacting protein, binding protein 1 (liprin beta 1)
	<u> </u>	Alternate: KIAA1230 protein
		A Monate. REPUTIZED DIOLOM
NM_030693	F:2.02 (7to19)	activating transcription factor 5
- NP 109618.1		
111 107010.1		
NM_008280	F:2.02 (5to19)	
- NP 032306.1		lipase
111 052500.1	· · · · · · · · · · · · · · · · · · ·	Subclass: lipase C precursor
<u></u>		Subclass: hepatic lipase
		Subclass: endothelial lipase precursor; endothelial cell-derived lipase
		Subclass: lipoprotein lipase precursor
		Subclass: Similar to lipoprotein lipase
NM_008407	F:2.02 (5to19)	
NP 032433.1]·	inter-alpha-trypsin inhibitor
		Subclass: pre-alpha (globulin) inhībitor, H3 polypeptide; Inter-alpha

		3 86
		Subclass: Inter-alpha-trypsin inhibitor heavy chain H1 precursor (ITI heavy
		chain H1) (Inter-alpha-inhibitor heavy chain 1) (Inter-alpha-trypsin inhibitor
		complex component III) (Serum-derived hyaluronan-associated protein)
		(SHAP)
	<u> </u>	Subclass: inter-alpha-trypsin inhibitor C-terminal
		Subclass: Inter-alpha-trypsin inhibitor heavy chain H2 precursor (ITI heavy
		chain H2) (Inter-alpha-inhibitor heavy chain 2) (Inter-alpha-trypsin inhibitor
		complex component II) (Serum-derived hyaluronan-associated protein)
		(SHAP)
'		Subclass: Inter-alpha-trypsin inhibitor heavy chain H4 precursor (ITI heavy
		chain H4) (Inter-alpha-inhibitor heavy chain 4) (Inter-alpha-trypsin inhibitor
		family heavy chain-related protein) (IHRP) (Plasma kallikrein sensitive
		glycoprotein 120) (PK-120) (GP120) (PRO1851) [Contains: GP57]
NM_009254	F:2.02 (5to19)	
. .	1 .2.02 (31013)	aning (an avertaine) and tainess inhibitor, slade P (evelbyrnin)
NP_033280.1	 	serine (or cysteine) proteinase inhibitor, clade B (ovalbumin) Subclass: serine (or cysteine) proteinase inhibitor, clade B (ovalbumin),
		member 6; protease in hibitor 6 (placental thrombin inhibitor
ļ		Subclass: serine (or cysteine) proteinase inhibitor, clade B (ovalbumin),
		member 3; squamous cell carcinoma antigen 1
	<u> </u>	Subclass: serine (or cysteine) proteinase inhibitor, clade B (ovalbumin),
		member 8; protease in hibitor 8 (ovalbumin type)
		Subclass: serine (or cysteine) proteinase inhibitor, clade B (ovalbumin),
		member 9; protease in hibitor 9 (ovalbumin type)
	1	Subclass: serine (or cysteine) proteinase inhibitor, clade B (ovalbumin),
•		member 1; protease in hibitor 2 (anti-elastase), monocyte/neutrophil; protease
		inhibitor 2 (anti-elastase), monocyte/neutrophil derived
,		Subclass: serine (or cysteine) proteinase inhibitor, clade B (ovalbumin),
		member 2; plasminogen activator inhibitor, type II (arginine-serpin)
		Subclass: serine (or cysteine) proteinase inhibitor, clade B (ovalbumin),
		member 10; protease inhibitor 10 (ovalbumin type, bomapin)
		Subclass: Similar to serine (or cysteine) proteinase inhibitor, clade B
		(ovalbumin), member 2 Subclass: Similar to serine (or cysteine) proteinase inhibitor, clade B
	ļ	(ovalbumin), member 8 Subclass: leupin
	 	Subclass: similar to Squamous cell carcinoma antigen 2 (SCCA-2) (Leupin
		Subclass: plasminogen activator inhibitor
		Subclass: hurpin
NTM 000659	F-2 02 (5+=10)	aldo-keto reductase
MM_009658	μ (3το 19)	aldo-keto reductase
NP 033788.1		Subclass: aldo-keto reductase family 1, member B1; aldehyde reductase 1;
	}	aldose reductase; low Km aldose reductase; Lii5-2 CTCL tumor antigen
	 	Subclass: aldo-keto reductase family 1, member B10; aldose reductase-like
		1; aldo-keto reductase family 1, member B11 (aldose reductase-like); aldose
	<u> </u>	11, and o reto reductase rating 1, member 111 (aldese reductase inte), aldese

40.

_	387
	reductase-like peptide; aldose reductase-related protein; small intestine
	reductase
	Subclass: similar to aldo-keto reductase family 1, member B10 (aldose
	reductase); aldose reductase-like 1; aldo-keto reductase family 1, member
1	B11 (aldose reductase-like
	Subclass: aldo-keto reductase family 1, member A1; aldehyde reductase;
	alcohol dehydrogenase
·	Subclass: aldo-keto reductase family 1, member D1; steroid-5-beta-
	reductase, beta polypeptide 1 (3-oxo-5 beta-steroid delta 4-dehydrogenase
	beta 1); steroid 5-beta-recluctase
	Subclass: aldo-keto reductase family 1, member C4 (chlordecone reductase;
	3-alpha hydroxysteroid dehydrogenase, type I; dihydrodiol dehydrogenase
	4); chlordecone reductase; type I 3-alpha-HSD; Chlordecone reductase Subclass: aldo-keto reductase family 1, member C1; dihydrodiol
	dehydrogenase 1; dihydrodiol dehydrogenase isoform DD1; type II 3-alpha-
	hydroxysteroid dehydrogenase; trans-1,2-dihydrobenzene-1,2-diol
	dehydrogenase; chlordecone reductase homolog; 20 alpha-hydroxysteroid
	dehydrogenase; aldo-keto reductase C; hepatic dihydrodiol dehydrogenase
	Subclass: similar to aldo-keto reductase
F-2 02 (5to19)	complement component
1 .2.02 (5.015)	complement component
 	Subclass: complement component 2 precursor; C3/C5 convertase
	Subclass: complement factor B preproprotein; B-factor, properdin; C3
	proactivator; C3 proaccelerator; glycine-rich beta-glycoprotein; C3/C5
	convertase
	Subclass: Similar to complement component 2
F:2.02 (5to19)	hypothetical protein BC0 1 3995
	Alternate: similar to hypothetical protein BC013995 Alternate: unnamed protein product
F:2.02 (5to19)	putative tumor suppressor FUS2
F:2.02 (5to19)	
F:2.02 (5to19)	
	putative tumor suppressor FUS2
F:2.01 (Min)	putative tumor suppressor FUS2 Peroxidase Subclass: glutathione peroxidase 6
F:2.01 (Min)	putative tumor suppressor FUS2 Peroxidase
F:2.01 (Min) F:2.01 (7to19)	putative tumor suppressor FUS2 Peroxidase Subclass: glutathione peroxidase 6 mannosidase, alpha, class 2B, member 1; mannosidase, alpha B, lysosomal
F:2.01 (Min) F:2.01 (7to19)	putative tumor suppressor FUS2 Peroxidase Subclass: glutathione peroxidase 6 mannosidase, alpha, class 2B, member 1; mannosidase, alpha B, lysosomal Alternate: Similar to mannosidase, alpha, class 2B, member 1; mannosidase,
F:2.01 (Min) F:2.01 (7to19)	putative tumor suppressor FUS2 Peroxidase Subclass: glutathione peroxidase 6 mannosidase, alpha, class 2B, member 1; mannosidase, alpha B, lysosomal
F:2.01 (Min) F:2.01 (7to19)	putative tumor suppressor FUS2 Peroxidase Subclass: glutathione peroxidase 6 mannosidase, alpha, class 2B, member 1; mannosidase, alpha B, lysosomal Alternate: Similar to mannosidase, alpha, class 2B, member 1; mannosidase,

2,5

		388
		Alternate: Unknown (protein for MGC:21066)
BC006621	F-2 01 (5to11)	KIAA0907 protein
	1.2.01 (5.011)	LENGTO PROCESS
AAH06621.1		Alternate: Similar to KIAA0907 protein
		Alichate. Similar to KiAA0507 protein
AB003502	F:2.01 (5to19)	G1 to S phase transition 1
BAA32526.1		
B74132320.1		Alternate: G1 to S phase transition 2
		Alternate: peptide chain release factor 3
		Alternate: similar to peptide chain release factor 3
		Alternate: polypeptide chain release factor 3b
		Alternate: KIAA1038 protein Alternate: unnamed protein product
	<u> </u>	Antemate: dimaned protein product
AK003237	F:2.01 (5to19)	hypothetical protein IMAGE3455200
BAB22661.1		
3123200111		
NM_008124	F:2.01 (5to19)	
NP 032150.2		gap junction protein
	· · · · · · · · · · · · · · · · · · ·	Subclass: gap junction protein, beta 1, 32kDa (connexin 32, Charcot-Marie
		Tooth neuropathy, X-linked); Gap junction protein, beta-1, 32kD (connexis
		32); gap junction protein, beta 1, 32kD (connexin 32, Charcot-Marie-Tooth
J		
		neuropathy, X-linked) Subclass: gap junction protein, beta 2, 26kDa (connexin 26); gap junction
	1	
		protein, beta 2, 26kD (connexin 26)
	1	Subclass: gap junction protein, beta 6 (connexin 30) Subclass: similar to Gap junction beta-4 protein (Connexin 30.3) (Cx30.3)
	:	Subclass: gap junction protein, beta 5 (connexin 31.1)
		Subclass: gap junction protein, beta 3, 31kDa (connexin 31); gap junction
		protein, beta 3, 31kD (connexin 31)
		Subclass: connexin25
		Subclass: similar to Gap junction beta-1 protein (Connexin 32) (Cx32)
•		(GAP junction 28 kDa liver protein)
		Subclass: gap junction protein, alpha 8, 50kDa (connexin 50); gap junction
		membrane channel protein alpha-8; connexin 50; Gap junction membrane
		channel protein alpha-8 (connexin 50); gap junction protein, alpha 8, 50kD
		(connexin 50) Subclass: gap junction protein, alpha 3, 46kDa (connexin 46); gap junction
		protein, alpha 3, 46kD (connexin 46)
•		Subclass: connexin 43; gap junction protein, alpha 1, 43kD (connexin 43);
		gap junction protein, alpha 1, 43kD
TM 000723	E-2 01 (5to 10)	nucleophosmin/nucle oplasmin 3; nucleoplasmin-3;
NM_008723		
		nucleophosmin/nucle oplasmin family, member 3
NP_032749.1		
		cathepsin Z; CTSZ

		209
M62361	F:2 (5to19)	sterol carrier protein 2
AAA40099.1		
NM_008331	F-2 (5to 10)	Tiple over (costs for MCC 14710)
_	F:2 (5to19)	Unknown (protein for MGC: 14710)
NP_032357.1	 	
		Alternate: interferon-induced protein
		Subclass: interferon—induced protein with tetratricopeptide repeats 1;
		Interferon, alpha-inducible protein (MW 56kD); interferon-induced protein
		56
		Subclass: retinoic acid- and interferon-inducible protein (58kD)
		Subclass: similar to Interferon-induced protein with tetratricopeptide repea
·		2 (IFIT-2) (Interferon-induced 54 kDa protein) (IFI-54K) (ISG-54 K)
		Subclass: interferon-induced protein with tetratricopeptide repeats 4
		Subclass: similar to Interferon-induced protein with tetratricopeptide repeat
		4 (IFIT-4) (Interferon-induced 60 kDa protein) (IFI-60K) (ISG-60) (CIG49)
		(Retinoic acid-induced gene G protein) (RIG-G)

	Mouse Gene	Behavio	r Description
	Protein		
	NM_007702	U:52.77	
	NP 031728.1	(YtoO)	coll dooth activates
ı	NF_U31/26.1	(100)	cell death activator Subclass: Cell death activator CIDE-A
	NM 007822	U:18.8	cytochrome P450,
-	NP_031848.1	(5to7)	cytochionic 1430,
		-	Subclass: cytochrome P450, subfamily IVA, polypeptide 11; fatty acid omega-
			hydroxylase; P450HL-omega; alkane-1 monooxygenase; lauric acid omega-
		1	hydroxylase (CYP4A11)
Ì			Subclass: cytochrome P450, subfamily IVB, polypeptide 1; cytochrome P450,
İ			subfamily IVB, member 1; microsomal monooxygenase
Ī	·		Subclass: cytochrome P450, subfamily IVF, polypeptide 2; leukotriene B4 omega
		1	hydroxylase; leukotriene-B4 20-monooxygenase
			Subclass: cytochrome P450, subfamily IVF, polypeptide 3; leukotriene B4 omega
Į	•		hydroxylase; leukotriene-B4 20-monooxygenase; cytochrome P450-LTB-omega
			Subclass: cytochrome P450, subfamily IVF, polypeptide 11
			Subclass: Cytochrome P450 4F12 (CYPIVF12)
			Subclass: cytochrome P450, subfamily IVF, polypeptide 8; microsomal
1			monooxygenase; flavoprotein-linked monooxygenase
-		<u> </u>	Subclass: similar to CYTOCHROME P450 4F6 (CYPIVF6)
f			Subclass: cytochrome P-450LTBV
	NM_008745	U:14.81	
	VP 032771.1	(YtoO)	Receptor tyrosine kinase
ſ			Subclass: neurotrophin receptor tyrosine kinase type 2
			Subclass: brain-derived neurotrophic factor receptor
Γ			Subclass: neurotrophin receptor trkC
			Subclass: TRKA
			Subclass: High affinity nerve growth factor receptor precursor (TRK1 transforming
-			tyrosine kinase protein) (p 140-TrkA) (Trk-A).
	NM_026574	U:12.76	KIAA1259 protein
E	NP_080850.1	(5to11)	
			Alternate: unnamed protein product
\vdash			Alternate: hypothetical protein DKFZp434B0616.1 - human
-	JM_021456	U:10.66	
1	_		Carboxylegterace
P	IP 067431.1	· · · · · · · · · · · · · · · · · · ·	Carboxylesterase Subclass: carboxylesterase 1 (monocyte/macrophage serine esterase 1); liver
ļ		1 0	carboxylesterase; carboxylesterase 2 (liver)

	1	1	Subclass: brain carboxylesterase hBr1
			Subclass: brain carboxylesterase hBr2
		1	Subclass: brain carboxylesterase hBr3
			Subclass: acyl coenzyme A:cholesterol acyltransferase
5			Subclass: egasyn
			Subclass: serine esterase N-terminal truncated (503 AA)
			Subclass: carboxylesterase hCE-2
	NM_013641	U:8.87	
10	NP_038669.1	(YtoM)	prostaglandin E receptor
			Subclass: prostaglandin E receptor, subtype EP1
		<u> </u>	
	AK004768	U:7.6	
	BAB23547.1	(YtoO)	oxysterol-binding protein-related protein
			Subclass: oxystero1-binding protein-like protein 3 isoform b; oxystero1-binding
15			protein-related protein 3; ysterol-binding protein 3; OSBP-related protein 3
			Subclass: oxystero1 binding protein-related protein 3 isoform 1a
			Subclass: oxystero1 binding protein-related protein 3 isoform 1d
			Subclass: oxystero1-binding protein-like protein 3 isoform c; oxystero1-binding
		ļ	protein-related protein 3; oxysterol-binding protein 3; OSBP-related protein 3
		İ	Subclass: oxystero1-binding protein-like protein 3 isoform e;oxysterol-binding
		<u> </u>	protein-related protein 3; oxysterol-binding protein 3; OSBP-related protein 3 Subclass: oxysterol-binding protein-like protein 3 isoform f;oxysterol-binding
20			protein-related protein 3; oxysterol-binding protein 3; OSBP-related protein 3
		<u> </u>	Subclass: oxystero1 binding protein-related protein 3 isoform 2c
			Subclass: oxystero1 binding protein-related protein 3 isoform 2d
	1		Subclass: oxystero1-binding protein-like 1A isoform B; oxysterol-binding protein-
			related protein 1; oxysterol-binding protein-like 1B; OSBP-related protein 1
			Subclass: oxysterol-binding protein-like 1A isoform C; oxysterol-binding protein-
			related protein 1; oxysterol-binding protein-like 1B; OSBP-related protein 1
25			Subclass: oxystero1-binding protein-like protein OSBPL1A
			Subclass: Oxystero 1-binding protein 2 (Oxysterol binding protein-related protein 4)
			(OSBP-related protein 4) (ORP-4).
		ļ ·	Subclass: oxystero1-binding protein-like protein 6 isoform b;oxysterol-binding
			protein-related protein 6; OSBP-related protein 6
		<u> </u>	Subclass: OSBP-related protein 7; ORP7
		<u> </u>	
30	AK011986	U:6.54	
	BAB27959.1	(7to19)	hypothetical protein FLJ32191
			Alternate: zinc finger protein 25
	NM 020568	U:6.5	
35	NP 065593.1	(YtoO)	KIAA1881 protein
22	MIE 003393.1	(1100)	KIATIOI PIOICIII
		.ii	

	1	1 .	392
	NM_013459	U:6.09	
	NP_038487.1	(5to11)	
		-	Complement factor D Subclass: Complement factor D precursor (C3 convertase activator) (Properdin
	İ		
			factor D) (Adipsin) Subclass: Chain, Mutant Of Factor D With Enhanced Catalytic Activity
-		1	Subclass: Chain, Human Complement Factor D In Complex With Isatoic
5	<u> </u>		
	ļ	- -	Anhydride Inhibitor
	NM_008182	U:5.76	glutathione trans ferase
	_		Standinone dans rotase
	NP_032208.1	(5to19)	
			Subclass: glutathione S-transferase A1; GST, class alpha, 1; glutathione S-
•	1		alkyltransferase A1; glutathione S-aryltransferase A1; S-(hydroxyalkyl)glutathione
	ļ		lyase A1; glutathione S-aralkyltransferase A1; GST-epsilon; glutathione S-
		1	transferase 2
10		1	Subclass: TPA: glutathione transferase A5
			Subclass: Chain A, Glutathione S-Transferase A1-1 (E.C.2.5.1.18)
			Subclass: Glutathione S-transferase A3-3 (GST class-alpha)
			Subclass: glutathione S-transferase A3
			Subclass: glutathione S-transferase A2; glutathione S-transferase 2; GST, class
			alpha, 2; liver GST2; glutathione S-alkyltransferase A2; glutathione S-aryltransferase
			A2; S-(hydroxyalkyl)glutathione lyase A2; glutathione S-aralkyltransferase A2;
			GST-gamma, HA subunit 2
15			JOS SHIPE STATE OF THE STATE OF
•	NM_009381	U:5.69	
	NP 033407.1	(YtoO)	Spot14 protein
	AK016553	U:5.55	
	i		
20	BAB30300.1	(YtoO)	heat shock transcription factor 2 binding protein; heat shock factor 2 binding protein
	U89406	U:5.43	
		l.	
	AAC36513.1	(YtoO)	fatty acid synthase
		 	
25	NM_025541	U:5.13	
	NP_079817.1	(YtoM)	HSPC146
			Alternate: dJ329 L24.2 (hypothetical 23.0 KD protein.)
	•		`
	AF281045	U:4.86	ribonuclease L (2',5'-oligoisoadenylate synthetase-dependent); ribonuclease 4
30	AAG33708.1	(5to11)	
			Alternate: A45771
	AK006096	U:4.75	
	BAB24407.1	(YtoO)	Similar to RIKEN cDNA 1700018018 gene
35		<u> </u>	

	•	393
NM_008495	U:4.6	beta-galactosidase binding lectin precursor; Lectin, galactose-binding, soluble, 1;
NP_032521.1	(7to11)	
NM 025429	U:4.44	
NP_079705.1	(5to19)	
	(3.012)	serine (or cysteine) proteinase inhibitor
	j	Subclass: serine (or cysteine) proteinase inhibitor, clade B (ovalbumin), member 1;
	1	protease inhibitor 2 (anti-elastase), monocyte/neutrophil; protease inhibitor 2 (anti-
		elastase), monocyte/neutrophil derived
		Subclass: serine (or cysteine) proteinase inhibitor, clade B (ovalbumin), member 9;
	,	protease inhibitor 9 (ovalbumin type)
	1	Subclass: serine (or cysteine) proteinase inhibitor, clade B (ovalbumin), member 10
		protease inhibitor 10 (ovalbumin type, bomapin)
		Subclass: serine (or cysteine) proteinase inhibitor, clade B (ovalbumin), member 8;
		protease inhibitor 8 (ovalbumin type)
·	}	Subclass: serine (or cysteine) proteinase inhibitor, clade B (ovalbumin), member 6;
		protease inhibitor 6 (placental thrombin inhibitor)
AF332052	U:4.08	
AAK56081.1_	(YtoO)	ATP citrate lyase
AK018226	U:4.01	serine (or cysteine) proteinase inhibitor, clade B (ovalbumin)
• •		(c. c), c. c) protoznaci zamonoż, c. c. c (c ()
XP_181363.1	(5to19)	
]	Subclass: serine (or cysteine) proteinase inhibitor, clade B (ovalbumin), member 1;
		protease inhibitor 2 (anti-elastase), monocyte/neutrophil; protease inhibitor 2 (anti-
	1	elastase), monocyte/neutrophil derived
		Subclass: serine (or cysteine) proteinase inhibitor, clade B (ovalbumin), member 9;
		protease inhibitor 9 (ovalbumin type)
		Subclass: serine (or cysteine) proteinase inhibitor, clade B (ovalbumin), member 8;
		protease inhibitor 8 (ovalbumin type)
1		Subclass: serine (or cysteine) proteinase inhibitor, clade B (ovalbumin), member 10;
<u> </u>	<u> </u>	protease inhibitor 10 (ovalbumin type, bomapin)
		Subclass: serine (or cysteine) proteinase inhibitor, clade B (ovalbumin), member 6;
	<u> </u>	protease inhibitor 6 (placental thrombin inhibitor)
	 	
NM_010831	U:3.91	
NP 034961.1	(YtoO)	Serine/threonine protein kinase
		Subclass: SNF1 -like kinase
		Subclass: Ser/Thr protein kinase PAR-1A
		Subclass: Ser/Thr protein kinase PAR-1B alpha
		Subclass: MAP/microtubule affinity-regulating kinase like 1; MARK4
		serine/threonine protein kinase
		Subclass: MAP/microtubule affinity-regulating kinase 2 isoform a; ELKL motif
		kinase 1; ELKL motif kinase

			394
		ı	Subclass: MAP/microtubule affinity-regulating kinase 2 isoform b; ELKL motif
			kinase 1; ELKL motif kinase
			Subclass: MAP/microtubule affinity-regulating kinase 3 long isoform
]	Subclass: Cdc25C associated protein kinase C-TAK1
			Subclass: 5'-AMP-activated protein kinase, catalytic alpha-2 chain (AMPK alpha-2
		İ	chain).
5			Subclass: KIAA0781 protein
			Subclass: KIAA0999 protein
	NM_023499	U:3.72	
	NP_075988.1	(YtoO)	immunoglobulin lambda light chain
10			Subclass: immunoglobulin lambda light chain VLJ region
			Subclass: Ig lambda VI THO
			Subclass: Ig lambda chain (BJP-DIA)
			Subclass: Ig lambda,anti-Rh(c).
			Subclass: Ig lambda chain V region
15			
	NM_009255	U:3.6	similar to tropomyosin, fibroblast - human
	NP_033281.1	(5to19)	
			Alternate: Protease Inhibitor; Proteinase Inhibitor
			Subclass: Glia derived nexin precursor (GDN) (Protease nexin I) (PN-1) (Protease
			inhibitor 7)
20			Subclass: Plasminogen Activator Inhibitor-1; Chain: A; Synonym: Pai-1,
	1 .		Endotheli al Plasminogen Activator Inhibitor, Pai
			Subclass: serine (or cysteine) proteinase inhibitor, clade E (nexin, plasminogen
	-		activator inhibitor type 1), member 1; plasminogen activator inhibitor, type I;
			Synonym: Pai-1, Endothelial Plasminogen Activator Inhibitor, Pai
			Subclass: prebeta-migrating plasminogen activator inhibitor
			Subclass: Cleaved Substrate Variant Of Plasminogen Activator Inhibitor-1
			Subclass: Active Form Of Human Pai-1
25			Subclass: serine (or cysteine) proteinase inhibitor, clade I (neuroserpin), member 1;
			protease inhibitor 12 (neuroserpin)
			Subclass: serine (or cysteine) proteinase inhibitor, clade I (neuroserpin), member 1
			Subclass: protease inhibitor 14; pancpin
	NM_021468	U:3.58	
30	NP_067443.1	(MtoO)	UNC13 (C. elegans)-like; homolog of rat Munc13 (diacylglycerol-binding)
	NM_007643	U:3.57	CD36 antigen (collagen type I receptor, thrombospondin receptor); CD36 antigen
	NP_031669.1	(YtoO)	(collagen type I)
35	AK007293	U:3.56	KIAA1879 protein
	BAB24937.1	(5to11)	

	1	1	1
	AK008016	U:3.37	
	AK008016	(YtoO)	Similar to RIKEN cDNA 2010001M09 gene
	AK008010	(1100)	Similar to Repen Control 201000111107 gene
5	NM_013623	U:3.35	orosomucoid 1 precursor; Orosomucoid-1 (alpha-1-acid glycoprotein-1); alpha-1-
	NP 038651.1	(7to19)	acid glycoprotein 1
		<u> </u>	
	57.00000	77.2.25	
	NM_020277	U:3.35	transierat receptor potential cation channel
	NP_064673.1	(5to11)	
10			Subclass: transient receptor potential cation channel, subfamily M, member 5;
			MLSN1 and TRP-related; MLSN1- and TRP-related; LTRPC5 protein
			Subclass: transient receptor potential cation channel, subfamily M, member 4
			Subclass: transient receptor potential-related channel 7, a novel putative Ca2+
			channel protein
	`		Subclass: transient receptor potential cation channel, subfamily M, member 2;
			transient receptor potential-related channel 7, a novel putative Ca2+ channel protein;
			transient receptor potential channel 7
			Subclass: transient receptor potential cation channel, subfamily M, member 8
15			Subclass: transient receptor potential cation channel, subfamily M, member 6
			Subclass: transient receptor potential cation channel, subfamily M, member 1;
			melastatin 1 [
	·	ļ	Subclass: TRP-related cation influx channer
		_	Subclass: channel-kinase 1 Subclass: similar to LTRPC7
		-	Alternate: melastatin 1
20		 	Alternate: melastam 1
	NM_007809	U:3.27	·
	NP_031835.1	(YtoO)	cytochrome P-450
		<u> </u>	Subclass: steroid 17alpha-monooxygenase (EC 1.14.99.9) cytochrome P450 17
			Subclass: aryl hydrocarbon (benzo[a]pyrene) hydroxylase (EC 1.14.14)
25		 	cytochrome P450 1A1
	J00544	U:3.24	immuno globulin J polypeptide, linker protein for immunoglobulin alpha and mu
	AAA38673.1	(YtoO)	polypeptides
	AAA30073.1	1100)	polypop tides
30	NM_031494	U:3.21	
<i>5</i> 0	1 -		7i for convertein
	NP_113682.1	(YtoM)	Zinc finger protein
	NM_008161	U:3.13	
	_	1	glutathione peroxidase
3 F	NP 032187.2	(YtoO)	Subclass: glutathione peroxidase 3 (EC 1.11.1.9)
3.5		 -	Subclass: glutatione peroxidase 3 (EC 1.11.1.9) Subclass: extracellular glutathione peroxidase
		-	Subclass: extracential gittathione peroxidase Subclass: glutathione peroxidase type 5 (GPX5)
	L		publiss. giliatinoite peroxidase type 5 (GI A5)

	_	_	396
			Subclass: plasma glutathione peroxidase
		,	
	NM_025724	U:3.12	
	NP_080000.1	(MtoO)	protein for MGC:26598
5			Alternate: Protein Similar to RIKEN cDNA 4921510H08 gene product
	ND 4 011125	U:3.1	phospholipid transfer protein
	NM_011125	ı	prosphoripid dansier protein
	NP_035255.1	(YtoO)	
			Alternate: dJ337O18.1.2 (Phospholipid Transfer Protein (Lipid Transfer Protein II)
•			(isoform 2))
10			Alternate: Similar to phospholipid transfer protein
	NM 012006	U:3.07	Peroxisomal acyl-coenzyme A thioester hydrolase 2 (Peroxisomal long-chain acyl-
	1 -		coA thioesterase 2) (ZAP128)
	NP_036136.1	(5to7)	
			A Iternate: peroxisomal long-chain acyl-coA thioesterase; peroxisomal long-chain
			acyl-coA thioesterase; putative protein
15			Alternate: Similar to peroxisomal long-chain acyl-coA thioesterase; peroxisomal
			long-chain acyl-coA thioesterase; putative protein
	NM_008361	U:3.05	interleukin 1, beta; preinterleukin 1 beta; interleukin 1; catabolin
	1 -	(5to7)	, oom , p.o.
	NP_032387.1	(3107)	
		<u> </u>	
20	NM_013559	U:2.97	
	NP_038587.1_	(YtoO)	heat shock protein
			Subclass: heat shock 105kD; heat shock 105kD alpha; heat shock 105kD beta
			Subclass: heat shock protein 70
			Subclass: similar to HEAT SHOCK 70 KDA PROTEIN 4 (HEAT SHOCK 70-
	, , , , , , , , , , , , , , , , , , , ,		RELATED PROTEIN APG-2) (HSP70RY)
25			Subclass: apg-2
	<u> </u>		Subclass: apg-1 Subclass: heat shock protein (hsp110 family)
			Subclass: HS24/P52
			S 400 (ass. 1152-1/1 52
30	AF127033	U:2.97	
	AAG02285.1	(YtoO)	fatty acid synthase; FAS
	NM 010062	U:2.89	deoxyribonuclease
	-	- 1	
	NP_034192.1	(5to11)	
35	·		Subclass: deoxyribonuclease II, lysosomal; DNase II, lysosomal
			Subclass: deoxyribonuclease II beta, isoform 1 precursor; DNase II-like a cid
			DNase; endonuclease DLAD
			<u> </u>

			327
	NM_011704	U:2.87	
	NP_035834.1	(5to7)	·
			Vanin
			Subclass: Vanin 1 (VNN1); pantetheinase
		_	Subclass: vanin 3 isoform 1; VNN3 protein; pantetheinase
5			Subclass: vanin 2, isoform 1; Vannin 2; pantetheinase
			Subclass: vanin 2, isoform 2; Vannin 2; pantetheinase
		_	Alternate: Biotinidase
		<u> </u>	
	AK018695	U:2.85	
0	NP 598527.1	(YtoM)	protein KIAA0143
0	141 390327.1	(11011)	Alternate; KIAA0953 protein
			interaction, and interest proton
	NM 011674	U:2.84	
_	-		Trong of our as
	NP_035804.1	(7to19)	Transferase Subclass: 2-hydroxyacylsphingosine 1-beta-galactosyltransferase precursor(UDP-
			galactose-ceramide galactosyltransferase) (Ceramide UDP-galactosyltransferase)
5			(Cerebroside synthase).
			Subclass: UDP glycosyltransferase 8 (UDP-galactose ceramide
			galactosyltransferase)
			Subclass: 2-hydroxyacylsphingosine 1-beta-galactosyltransferase (EC 2.4.1.45)
•			Subclass: UDP-glucuronosyltransferase 2B17 precursor, microsomal
			(UDPGT)(C19-steroid specific UDP-glucuronosyltransferase).
		_	Subclass: UDP-glucuronosyltransferase 2B15
			Subclass: UDP glycosyltransferase 2 family, polypeptide A1; UDP
0			glucuronosyltransferase 2 family, polypeptide A1
			Subclass: UDP glucuronosyltransferase 1A5
			Subclass: UDP glucuronosyltransferase 1A6
•		<u> </u>	Subclass: UDP-glucuronosyltransferase 1A7
			Subclass: UDP glucuronosyltransferase 1A8
5		<u> </u>	Subclass: UDP glycosyltransferase 1 family, polypeptide A9
		 	Subclass: UDP-glucuronosyltransferase 1A10
			Subclass: UDP-glucuronosyltransferase 1-2 precursor, microsornal (UDPGT)(UGT-
		<u> </u>	1B) (UGT1*2) (UGT1-02) (UGT1.2) (UGT1A2) (UGT1B) (HLUGP4).
		<u> </u>	Subclass: UDP-glucuronosyltransferase 2B
			Subclass: UDP glucuronosyltransferase 2B4
			Subclass: UDP-glucuronosyltransferase 2B7, microsomal (UDP GT)(3,4-catechol
0			estrogen specific) (UDPGTH-2).
			Subclass: UDP-glucuronosyltransferase (EC 2.4.1,-) 2B-10
	NM 023455	U:2.75	putative N-acetyltransferase Camello 2
	1 -	1	
	NP_075944.1	(5to19)	, Ma
5			Alternate: N-acetyltransferase 8; kidney- and liver-specific gene product; kidney-
			and liver-specific gene

	1 .	1	Alternate: GLA
			Alternate: hypothetical protein TSC501 [imported]
	NM_023478	U:2.74	uroplakin 3
5	NP_075967.1	(5to19)	
	NM_016774	U:2.73	• .
	NP 058054.1	(YtoM)	ATP synthase
			Subclass: ATP synthase, H+ transporting, mitochondrial F1 complex, beta
			polypeptide; ATP synthase, H+ transporting, mitochondrial F1 complex, beta
10	NM_011146	U:2.68	
	NP 035276.1	(5to11)	
	NP_033276.1	(3.011)	peroxisome proliferative activated receptor
			Subclass: peroxisome proliferative activated receptor gamma
			Subclass: peroxisome proliferative activated receptor gamma, isoform 2; PPAR-
			gamma; peroxisome proliferator activated receptor gamma
5			Subclass: peroxisome proliferative activated receptor gamma, isoform 1; PPAR-
		· · · · ·	gamma; peroxisome proliferator activated receptor gamma
	NM_013771	U:2.67	· ·
	NP_038799.1	(YtoM)	Metalloprotease
		1	Subclass: ATP-dependent metalloprotease YME1L
20			Subclass: YME1-like 1 isoform 1; ATP-dependent metalloprote ase FtsH1 homolog
	AK002979	U:2.67	calcyon
	BAB22492.1	(5to19)	
_			
5	AK005609	U:2.62	
	BAB24148.1	(5to19)	similar to RIKEN cDNA 1700001L19 [Mus musculus]
	X03796	U:2.61	
	CAA27422.1	(YtoM)	Aldolase
0	•		Subclass: aldolase C
			Subclass: aldolase A
			Subclass: aldolase B
		<u> </u>	Subclass: Fructose 1,6-Bisphosphate Aldolase From Human Liver
5	NM 019415	U:2.6	
	NP 062288.1	(5to11)	
		(2.011)	So lute carrier family 12
			Subclass: Solute carrier family 12 member 3 (Thiazide-sensitive sodium-chloride
			cotransporter) (Na-Cl symporter)
			Subclass: solute carrier family 12 (sodium/potassium/chloride transporters), member
		<u></u>	2; Solute carrier family 12 (sodium/potassium/chloride transporters),

1	ı	399 Subclass: solute carrier family 12 (potassium/chloride transporters), member 7;
	1	
	-	potassium/chloride transporter KCC4 Subclass: solute carrier family 12, (potassium-chloride transporter) member 5
	†	oucouron source current running 125, (potabolium emotitus transporter) member 3
AK009937	U:2.57	
BAB26596.1	(YtoM)	hypothetical protein FLJ12118
		Alternate: Synthetase
		Subclass: cysteine-tRNA ligase, isoform a; cysteine translase; cysteine-tRNA
	 	synthetase
		Subclass: cysteine-tRNA ligase, isoform b; cysteine translase; cysteine-tRNA
,	 	synthetase
NM_023137	U:2.56	
NP 075626.1	(YtoO)	ubiquitin-like protein FAT10
AK015750	U:2.56	
BAB29956.1	(YtoO)	Sulfotransferase
		Subclass: sulfotransferase, estrogen-preferring
		Subclass: thyroid hormone sulfotransferase
		Subclass: sulfotransferase family 1A
		Subclass: sulfotransferase family, cytosolic, 1A, phenol-preferring, member 1
		Subclass: SULT1C sulfotransferase; sulfotransferase family, cytosolic, 1C, mem
		C2
		Subclass: sulfotransferase 1C1
		Subclass: Phenol-sulfating phenol sulfotransferase 1 (P-PST) (Thermostable phenol
		sulfotransferase) (Ts-PST) (HAST1/HAST2) (ST1A3).
·		Subclass: phenol-preferring phenol sulfotransferase 2
		Subclass: aryl sulfotransferase ST1A2
		Subclass: aryl sulfotransferase ST1A3
		Subclass: arylamine sulfotransferase
		Subclass: alcohol/hydroxysteroid sulfotransferase; hSTa
		Subclass: sulfotransferase family, cytosolic, 2A, dehydroepiandrosterone(DHEA
		preferring, member 1; sulfotransferase family 2A, dehydroepiandrosterone (DHEA
		preferring, member 1
		Subclass: hydroxysteroid sulfotransferase SULT2B1a
		Subclass: hydroxysteroid sulfotransferase SULT2B1b
1 Tr 0.0 c c c c	71.0 ==	
AK002693	U:2.55	
BAB22288.1	(YtoO)	diacylglycerol acyltransferase
		Subclass: diacylglycerol O-acyltransferase 2 like 1; iacylglycerolacyltransferase 2
		like
		Subclass: diacylglycerol acyltransferase 2
1		
VM 011214	U:2.54	

Ì		Subclass: protein tyrosine phosphatase, receptor type, U isoform 1 precursor;
		protein tyrosine phosphatase J; protein tyrosine phosphatase receptor omicron
		Subclass: protein tyrosine phosphatase, receptor type, U isoform 2 precursor;
		protein tyrosine phosphatase J; protein tyrosine phosphatase receptor omicron
		Subclass: protein tyrosine phosphatase, receptor type, U isoform 3
		precursor;protein tyrosine phosphatase J; protein tyrosine phosphatase receptor
		omicron
İ		Subclass: protein tyrosine phosphatase receptor omicron
5		Subclass: receptor phosphatase PCP-2
		Subclass: FMI protein
		Subclass: protein tyrosine phosphatase, receptor type, K precursor; protein-tyrosine
	·	phosphatase, receptor type, kappa; protein-tyrosine phosphatase kappa; protein-
ļ		tyrosine phosphatase kappa precursor
		Subclass: protein tyrosine phosphatase, receptor type, M precursor; protein tyrosine
		phosphatase, receptor type, mu polypeptide; protein tyrosime phosphatase mu
		precursor
		Subclass: protein tyrosine phosphatase, receptor type, T, isoform 2 precursor;
		receptor protein tyrosine phosphatase
0 [·	Subclass: protein tyrosine phosphatase sigma
	·	Subclass: protein tyrosine phosphatase, receptor type, sigrma, isoform 1 precursor;
1		protein tyrosine phosphatase PTPsigma
Ī		Subclass: protein tyrosine phosphatase, receptor type, sigrna, isoform 2 precursor;
	·	protein tyrosine phosphatase PTPsigma
		Subclass: protein tyrosine phosphatase, receptor type, sigrna, isoform 3 precursor;
1		protein tyrosine phosphatase PTPsigma
Ī		Subclass: protein tyrosine phosphatase, receptor type, sigrma, isoform 4 precursor;
1		protein tyrosine phosphatase PTPsigma
5		Subclass: PTPsigma-(brain)
Ī		Subclass: protein tyrosine phosphatase delta
Ī		Subclass: protein tyrosine phosphatase, receptor type, D isoform 4 precursor;
!		protein tyrosine phosphatase, receptor type, delta polypeptide; protein tyrosine
		phosphatase delta
Ī		Subclass: protein tyrosine phosphatase, receptor type, D isoform 2 precursor;
	İ	protein tyrosine phosphatase, receptor type, delta polypeptide; protein tyrosine
		phosphatase delta
-		Subclass: protein tyrosine phosphatase, receptor type, D is oform 3 precursor;
		protein tyrosine phosphatase, receptor type, delta polypeptide; protein tyrosine
	·	
-		phosphatase delta
		Subclass: protein tyrosine phosphatase, receptor type, F, isoform 2 precursor;
		protein tyrosine phosphatase, receptor type, F polypeptide; receptor-linked protein-
		tyrosine phosphatase LAR; Leukocyte antigen-related tyrosine phosphatase; LCA-
o		homolog

	NM_019935	U:2.52	OVO-like 1 binding protein; putative transcription factor OVO-like 1; ovo
	NP_064319.1	(5to11)	(Drosophila) homolog-like 1
			Alternate: hypothetical protein, similar to (AF134804) putative zinc finger
			transcription factor OVO1
			Alternate: zinc finger protein 339; ovo-like 2 (Drosophila)
5			
	NM_033174	U: 2. 51	
	NP 149409.1	(YtoO)	snRNP
			Subclass: snRNP polypeptide B.
_0	NM_008714	U:2.5	Neurogenic locus notch homolog protein (Notch)
. 0	_ ·	l l	reducing the rotal notion of protein (reducin)
	NP_032740.1	(5to19)	
			Subclass: Neurogenic locus notch homolog protein 1 precursor (Notch 1) (hN1)
			(Translocation-associated notch protein TAN-1)
			Subclass: NOTCH2 protein
			Subclass: Notch3
.5 .			Subclass: Notch homolog 4 (Drosophila); Notch, drosophila, homolog of, 4; Notch
			(Drosophila) homolog 4
	·		Alternate: transmembrane protein Jagged 1
			Alternate: fibrillin
			Subclass: fibrillin 1; Fibrillin-1
			Subclass: fibrillin 2
20			Subclass: similar to fibrillin
	D. C. 010000	15	
	NM_019992	U:2.47	
	NP_064376.1	(YtoO)	BCR downstream signaling 1
5	NM 019640	U:2.47	
_	NP 062614.1	!	Phosphatidylinositol transfer protein
	NF 002014.1	(1101/1)	Subclass: dJ353E16.1 (phosphatidylinositol transfer protein beta)
	 	 	Subclass: Phosphatidylinositol transfer protein alpha isoform (PtdIns transfer
		.].	
			protein alpha) (PtdInsTP) (PI-TP-alpha).
^		 	Subclass: NIR2
0		<u> </u>	Subclass: homologue of Drosphila retinal degeneration B gene
		1	Subclass: PYK2 N-terminal domain-interacting recept or 3; KIAA1457 protein;
			likely ortholog of mouse retinal degeneration B2 homolog (Drosophila) (Rdgb2)
	NM_007824	U:2.47	
	_		nutochromo D450
	NP 031850.1		cytochrome P450 Subclass: cytochrome P450, subfamily VIIA, polypepti de 1; cholesterol 7-alpha-
_			•
5		1	hydroxylase; cholesterol 7 alpha-monooxygenase
		<u> </u>	Subclass: sterol 12-alpha-hydroxylase CYP8B1

402

		402
	-{	Alternate: prostacyclin synthase
NM 011076	U:2.45	
NP 035206.1	(YtoO)	P glycoprotein.
14F_035200.1	(1100)	Subclass: ATP-binding cassette, sub-family B (MDR/TAP), member 1; P
		glycoprotein 1/multiple drug resistance 1; P-glycoprotein-1/multiple drug resistan
		1; multidrug resistance 1
	 	Subclass: ATP-binding cassette, subfamily B, member 4 isoform B; P glycoprote
		3/multiple drug resistance 3; P-glycoprotein-3/multiple drug resistance-3; multiple
		drug resistance 3
		Subclass: ATP-binding cassette, subfamily B, member 4 isoform C; P glycoprote
		3/multiple drug resistance 3; P-glycoprotein-3/multiple drug resistance-3; multiple
		drug resistance 3
	 	Subclass: ATP-binding cassette, sub-family B (MDR/TAP), member 11; ABC
		member 16, MDR/TAP subfamily; progressive familial intrahepatic cholestasis 2;
		bile salt export pump
		Alternate: bile salt export pump
		· · · · · · · · · · · · · · · · · · ·
NM_009345	U:2.43	
_		deoxynucleotidyltransferase, terminal; Terminal deoxynucleotidyltransferase
NP 033371.1	(YtoO)	Alternate: polymerase (DNA directed), mu; polymerase (DNA-directed), mu
AK009815	U:2.42	
BAB26519.1	(YtoO)	1,4-alpha-glucan branching enzyme (EC 2.4.1.18)
NM_011178	U:2.42	
NP 035308.1	(YtoM)	Proteinase
141 033308.1	(1 (0)(1)	
	,	Subclass: Myeloblastin (Leukocyte proteinase 3) (PR-3) (PR3) (AGP7)(Wegene
	<u> </u>	autoantigen) (P29) (C-ANCA antigen) (Neutrophil proteinase 4) (NP-4).
		Subclass: proteinase 3
		Subclass: medullasin
		Subclass: Human Heparin Binding Protein
		Subclass: azurocidin Subclass: cationic antimicrobial protein CAP37
		Subclass: leukocyte elastase (EC 3.4.21.37)
	 	Substance (DO 5.4.21.37)
	U:2.42	forkhead box F2; forkhead (Drosophila)-like 6
NM_010225	U:2.42	
NM_010225 NP_034355.1	(5to11)	
	1	Alternate: transcription factor FREAC-2
	1	Alternate: transcription factor FREAC-2 Alternate: forkhead box F1; forkhead (Drosophila)-like 5; Forkhead, drosophila, homolog-like 5; forkhead-related activator 1

NM_007760	U:2.41	carnitine acetyltransferase
NP_031786.		
		Subclass: Carnitine O-acetyltransferase (Carnitine acetylase) (CAT)
		Subclass: carnitine acetyltransferase isoform 1
ļ		Subclass: carnitine acetyltransferase isoform 2
		Subclass: carnitine acetyltransferase isoform 3
NMI_010846	U:2.39	
NP 034976.1	(YtoO)	Interferon-induced protein
		Subclass: interferon-induced Mx protein
		Subclass: myxovirus resistance protein 1; inter feron inducible protein p78;
		interferon-regulated resistance GTP-binding protein
		Subclass: interferon-induced viral resistance protein MxB
		Subclass: MX2
		Alternate: dynamin
		Subclass: dynamin 2; Dynamin II
		Subclass: Dynamin 3 (Dynamin, testicular) (T-dynamin).
NM_008151	U:2.39	G protein-coupled receptor
_	1	O protein-coupled receptor
NP_032177.1	(7to19)	
·		Subclass: G protein-coupled receptor 12
		Subclass: G protein-coupled receptor 3; adenylate cyclase constitutive activator
		Subclass: G protein-coupled receptor 6
D00208	U:2.39	S100 calcium-binding protein A4; 18A2; 42A; S100 calcium-binding protein A4
BAA00148.1	(5to11)	(calcium protein, calvasculin, metastasin, murine placental homolog); malignant
BAA00146.1	(31011)	1
		transformation suppression 1
·	- 	<u> </u>
NM_012050	U:2.38	
NP 036180.1	(YtoO)	osteomodulin
NM_019748	U:2.38	
NP 062722.1	(YtoM)	SUMO-1 activating enzyme
		Subclass: SUMO-1 activating enzyme subunit 1
NM_008273	U:2.36	homeo box D11; homeo box 4F; Hox-4.6, mouse, homolog of; homeobox protein
NP_O32299.1	(5to11)	Hox-D11
NM_009676	U:2.36	Aldehyde oxidase
NP_O33806.1	(5to7)	
	<u> </u>	Subclass: aldehyde oxidase 1
L		Subcrass, aidenyde oxidase 1

	_	404
	_	Subclass: Similar to aldehyde oxidase 1
NM 009773	U:2.35	
NP 033 903.1	(MtoO)	mitotic checkpoint protein kinase
112 033 303.1	(Miss)	Subclass: MAD3-like protein kinase
		Subclass: budding uninhibited by benzimidazoles 1 beta
NM_03O127	U:2.32	
NP_084403.1	(YtoO)	serine protease
		Subclass: serine protease HTRA3
		Subclass: protease, serine, 11
		Subclass: serine protease HtrA2-p7
NM_011086	U:2.32	unnamed protein product
NP_035216.1	(5to11)	FF
141_055210.1	(3,011)	
		Alternate: FYVE finger-containing phosphoinositide kinase (1-phosphatidylinos
	-	4-phosphate 5-kinase) (PIP5K) (PtdIns(4)P-5-kinase) (p235)
		Alternate: similar to FYVE finger-containing phosphoinositide kinase (1-
ļ		phosphatidylinositol-4-phosphate kinase) (PIP5K) (PtdIns(4)P-5-kinase) (p235) Alternate: hypothetical protein MGC40423
		Alternate: hypothetical protein wide40423
V00795	U:2.3	
CAA241 76.1	(5to19)	Immunoglobulin heavy chain
		Subclass: immunoglobulin heavy chain constant region
	<u> </u>	Subclass: Ig gamma 2 H chain BUR
		Subclass: Ig gamma-2 chain C region
		Subclass: immunoglobulin gamma 2 heavy chain constant region
		Subclass: recombinant IgG1 heavy chain
	<u> </u> -	Subclass: immunoglobulin lambda heavy chain
	 	Subclass: Ig gamma-1 chain C region - human
NM_016922	U:2.29	
NP 058618.1	(YtoO)	Sulfotransferase
		Subclass: Galactosylceramide sulfotransferase (GalCer sulfotransferase)
·		(Cerebroside sulfotransferase) (3'-phosphoadenylylsul.fate: galactosylceramide 3'-
		sulfotransferase) (3'-phosphoadenosine-5'phosphosulfate: GalCer sulfotransferase
	<u> </u>	Subclass: glycoprotein beta-Gal 3'-sulfotransferase
		Subclass: beta-galactose-3-O-sulfotransferase 3
		Subclass: beta-galactose-3-O-sulfotransferase, 4
NM_013739	U:2.29	
NP 038767.1	1	hypothetical protein FLJ22570

	1		405
	NM_008673	U:2.29	
	NP O 32699.1	(MtoO)	N-acetyltransferase
			Subclass: arylamine N-acetyltransferase 1
5	NM_026189	U:2.29	KIAA1706 protein
	NP_080465.1	(5to11)	
			Alternate: similar to RIKEN cDNA 2310005P05
		- 	Alternate: unnamed protein product
		- 	Alternate: umianieu protein product
10	AF047725	U:2.28	
	AAD 13720.1	(5to11)	cytochrome P450, subfamily IIC
	. AAD 13720.1	(3(011)	Subclass: cytochrome P450, subfamily IIC (mephenytoin 4-hydroxylase),
			polypeptide 18; cytochrome P450, subfamily IIC (mephenytoin 4-hydroxylase),
		 	polypeptide 17; microsomal monooxygenase; flavoprotein-linked monooxygenase Subclass: cytochrome P450, subfamily IIC (mephenytoin 4-hydroxylase),
			polypeptide 19; mephenytoin 4'-hydroxylase; microsomal monooxygenase;
		-	xenobiotic monooxygenase; flavoprotein-linked monooxygenase Subclass: Cytochrome P450 2C8 (CYPIIC8) (P450 form 1) (P450 MP-12/MP-20)
15		 	(P450 IIC2) (S-mephenytoin 4-hydroxylase) Subclass: cytochrome P450, subfamily IIC, polypeptide 9; cytochrome P450,
יב			
			subfamily IIC (mephenytoin 4-hydroxylase), polypeptide 10; mephenytoin 4-
]		hydroxylase; microsomal monooxygenase; xenobiotic monooxygenase; flavoprotein-
			linked monooxygenase
			Subclass: Cytochrome P450 2C10 (CYPIIC10) (P450 MP-8) (S-mephenytoin 4-
		 	hydroxylase) (P-450MP)
	NM 058212	U:2.26	cerebellum D4
	NP_478119.1	(5to19)	
	4 /8119.1	(31019)	
20			
	NM_011897	U:2.25	·
	-	1	The state of the s
	NP_036027.1		antagonist of FGF and/or EGF signaling
		-	Subclass: sprouty (Drosophila) homolog 2
25		1	Subclass: Sprouty homolog 1 (Spry-1). Subclass: sprouty homolog 3; antagonist of FGF signaling
23			Subclass: sprouty homolog 5, antagonist of PGF stenaring
	NM_O18861	U:2.25	
	1 -	1 1	
	NP_061349.1		neutral amino acid transporter Subclass: solute carrier family 1, member 4; glutamate/neutral amino acid
		1 1	-
30		1	transporter; alanine/serine/cysteine/threonine transporter
J ()			Subclass: solute carrier family 1 (neutral amino acid transporter), member 5 Subclass: solute carrier family 1 (glial high affinity glutamate transporter), member
		ľĺ	3
	ı	1 . F	

		_	406
			Subclass: solute carrier family 1 (neuronal/epithelial high affinity glutamate
			transporter, system Xag), member 1
			Subclass: solute carrier family 1, member 2; H.sapiens mRNA for glutamate
			transporter; glutamate/aspartate transporter II; excitatory amino acid transporter 2;
			glial high affinity glutamate
			Subclass: solute carrier family 1 (glutamate transporter), member 7
			Subclass: neutral amino acid transporter B
5			Subclass: Na+-dependent neutral amino acid transporter SATT
			Subclass: sodium-dependent neutral amino acid transporter
			Subclass: Excitatory amino acid transporter 3 (Sodium-dependent
			glutamate/aspartate transporter 3) (Excitatory amino-acid carrier1) (Neuronal and
			epithelial glutamate transporter).
			Subclass: neuronal and epithelial glutamate transporter
			Subclass: retinal glutamate transporter EAAT5
10			<u>.</u>
	AF126834	U:2.24	
	AAD20642.1	(YtoO)	Intermediate filament binding protein
			Subclass: Periplakin (195 kDa cornified envelope precursor) (190 kDa
			paraneoplastic pemphigus antigen).
	`	•	Subclass: envoplakin
15			Subclass: Plectin 1 (PLTN) (PCN) (Hemidesmosomal protein 1) (HD1).
	İ		Subclass: bullous pemphigoid antigen 1, isoform 1eA precursor; dystonin;
			hemidesmosomal plaque protein
		į	Subclass: bullous pemphigoid antigen 1 isoform 1eB precursor; bullous pemphigoid
			antigen 1; bullous pemphigoid antigen 1 (230/240kD); dystonin; hemidesmosomal
			plaque protein
			Subclass: Bullous pemphigoid antigen 1, isoforms 6/9/10 (Trabeculin-beta) (Bullous
			pemphigoid antigen) (BPA) (Hemidesmosomal plaque protein) (Dystonia
			musculorum protein).
		\ \ \ \ \	Subclass: Bullous pemphigoid antigen 1 isoforms 1/2/3/4/5/8 (230 kDa bullous
,			pemphigoid antigen) (BPA) (Hemid esmosomal plaque protein) (Dystonia
			musculorum protein).
20			
į	U67189	U:2.23	
<u> </u> -	AAB50619.1	(YtoM)	regulator of G protein signalling 16
! !	NM_008762	U:2.23	olfactory receptor
25	NP_032788.1	(5to19)	
!		 	Subclass: olfactory receptor, family 2, subfamily C, member 1
;		1	Subclass: olfactory receptor, family 2, subfamily C, member 3
1			Subclass: similar to olfactory receptor, family 2, subfamily C, member 3
		 	Subclass: Olfactory receptor 2B6 (Hs6M1-32) (Olfactory receptor 6-31) (OR6-31)
20		 	Subclass: Olfactory receptor, family 2, subfamily B, member 2
30		1	Subclass: similar to Olfactory receptor 2B2 (Olfactory receptor 6-1) (OR6-1)
1			
1		<u></u>	(Hs6M1-10)

			407
			Subclass: olfactory receptor, family 2, subfamily H, member 3; Olfactory receptor
			Subclass: Olfactory receptor 2H2 (Hs6M1-12)
		-	Subclass: similar to Olfactory receptor 2H1 (Hs6M1-16) (Olfactory receptor 6-2)
			(OR6-2) (OLFR42A-9004.14/9026.2)
			Subclass: similar to Olfactory receptor 2J3 (Olfactory receptor 6-6) (OR6-6)
			(Hs6M1-3)
5		- 	Subclass: similar to Olfactory receptor 2B3 (Olfactory receptor 6-4) (OR6-4)
_			(Hs6M1-1)
			Subclass: olfactory receptor, family 5, subfamily V, member 1
			Subclass: olfactory receptor, family 2, subfamily W, member 1
		 	Subclass: olfactory receptor, family 2, subfamily J, member
	-	<u> </u>	Subclass: similar to olfactory receptor MOR256-3
)			Subclass: similar to olfactory receptor MOR256-12
			Subclass: similar to olfactory receptor MOR256-14
			Subclass: olfactory receptor 89
			Subclass: similar to olfactory receptor 89
			Subclass: similar to 573K1.15 (mm.17M1-6 (novel 7 transmembrane receptor
			(rhodopsin family) (olfactory receptor LIKE) protein))
5			Alternate: seven transmembrane helix receptor
			Alternate: similar to seven transmermbrane helix receptor
	NM 013746	U:2.22	pleckstrin homology domain containing, family B (evectins) member 1; PH domain
	NP 03 8774.1	(YtoO)	containing protein in retina 1; PH domain containing, retinal 1
)	14 05 07 74.1	(1100)	Containing protoin in rotals 1, 112 Contain Containing, Totals 1
	NM_009613	U:2.22	
	NP 033743.1	(MtoO)	Metalloprotease/disintegrin-like protein
	Ì		Subclass: ADAM 11 precursor (A disintegrin and metalloproteinase domain 11)
			(Metalloproteinase-like, disintegrin-like, and cysteine-richprotein)
			Subclass: a disintegrin and metalloproteinase domain 22 isoform 2 proprotein;
			MDC2 delta
			Subclass: a disintegrin and metalloproteinase domain 22 isoform 3 proprotein;
5			MDC2 delta
			Subclass: a disintegrin and metalloproteinase domain 22 isoform 4 proprotein;
			MDC2 delta
			Subclass: a disintegrin and metalloproteinase domain 22 isoform 5 proprotein;
			MDC2 delta
			Subclass: ADAM 22 (A disintegrin and metalloproteinase domain
			22)(Metalloproteinase-like, disintegr in-like, and cysteine-rich protein 2)
			(Metalloproteinase-disintegrin ADAM22-3).
		1	Subclass: a disintegrin and metalloproteinase domain 8
)		 	Subclass: a disintegrin and metalloproteinase domain 21 preproprotein
•			Subclass: a disintegrin and metallop totemase domain 21 preproprotein
		<u> </u>	Subclass: a disintegrin and metalloproteinase domain 28 isoform 1
		1	
		1	Subclass: disintegrin and metalloproteinase domain 19

408

			408
			Subclass: a disintegrin and metalloproteinase domain 19 isoform 1 preproprotein;
			meltrin beta
			Subclass: a disintegrin and metalloproteinase domain 19 isoform 2 preproprotein;
			meltrin beta
	·		Subclass: a disintegrin and metalloprotease domain 12 isoform 2 preproprotein; A
			disintegrin and metalloproteinas e domain 12(Meltrin-alpha, mouse, homolog of);
			meltrin alpha
			Subclass: a disintegrin and metalloproteinase domain 33 isoform alpha; disintegri
			and reprolysin metalloproteinase familyprotein; metalloprotease disintegrin
5			Subclass: a disintegrin and metalloproteinase domain 15 (metargidin)
			Subclass: meltrin-S
			Subclass: metalloprotease-disintegrin meltrin beta
-			
	NM_008008	U:2.22	fibroblast growth factor 7 precursor; keratinocyte growth factor
10	NP_032034.1	(5to19)	
			
	AF366393	U:2.21	
		1	
	AAK53703.1	(YtoM)	Protein phosphatase Subclass: protein phosphatase 2 (formerly 2A), regulatory subunit B (PR 52), beta
15		1 -	isoform
TD			Subclass: protein phosphatase 2.A BR gamma subunit
		_	Subclass: protein phosphatase 2.A1 B gamma subunit IMYPNO1
	7. 6.000107	77.0.0	
	NM_009127	U:2.2	
	NP 033153.1	(YtoM)	Desaturase (C) 1 (
		1	Subclass: Acyl-CoA desaturase (Stearoyl-CoA desaturase) (Fatty acid desaturase)
20			(Delta(9)-desaturase).
	NM_016894	U:2.2	receptor (calcitonin) activity modifying protein 1 precursor; calcitonin receptor-like
	· -	I	
	NP_058590.1	(5to11)	receptor activity modifying protein 1
0.5	DD 6 010560	11.0.0	
25	NM_013560	U:2.2	heat shock protein
	NP_038588.1	(5to7)	
			Subclass: heat shock 27kDa protein 1; heat shock 27kD protein 1
			Subclass: heat shock protein 27
		<u> </u>	Subclass: Unknown (protein for IMAGE:3906970)
30		1	Subclass: similar to Heat shock 27 kDa protein (HSP 27) (Stress-responsive protein
			27) (SRP27) (Estrogen-regulated 24 kDa protein) (28 kDa heat shock protein
		-	
	NM_019977	U:2.18	aldehyde reductase (aldose reductase) like 6; similar to mouse aldehyde reductase 6
	n 1111 111177//	IU.Z.10	pridory do reducido (ardose reducido) rice o, surmar to mouse aldenyde reducidse (
	NP 064361.1	(YtoO)	(renal); myo-inositol oxygenase; kidney-specific protein 32

			409
	NM_031194	U:2.18	
	NP 112471.1	(MtoO)	Organic anion transporter
			Subclass: organic anion transporter 3
		1	Subclass: solute carrier family 22 member 6 isoform b; renal organic anion
			transporter 1; para-aminohippurate transporter
			Subclass: solute carrier family 22 member 6 isoform a; renal organic anion
5			transporter 1; para-aminohippurate transporter
			Subclass: solute carrier family 22 member 6 isoform e; renal organic anion
			transporter 1; para-aminohippurate transporter
			Subclass: solute carrier family 22 member 7 isoform a; organic anion transporter 2;
			liver-specific transporter
			Subclass: solute carrier family 22 member 4; organic cation transporter 4; integral
			membrane transport protein
			Subclass: urate anion exchanger 1 is oform b; organic anion transporter 4-like; urate
			transporter 1; solute carrier family 22 member 12
0		<u> </u>	Subclass: solute carrier family 22 member 1 isoform a; organic cation transporter 1
			Subclass: RST
			Subclass: OAT4
		ļ	Subclass: organic anion transporter 2
		<u>. </u>	Subclass: renal organic anion transporter 1
5		<u> </u>	Subclass: hUST3
	ļ	 	Subclass: OCTN1
	<u> </u>	- }	Subclass: OCTN2
		 	Subclass: extraneuronal monoamine transporter
)		 	Subclass: bA288H12.2 (organic cation transporter, liver)
,			Subclass: organic cation transporter OKB1
	NM 020051	U:2.17	putative bHLH transcription factor
	NP 064435.1	(YtoO)	
	. 1 _004433.1	(1.00)	
			Alternate: Achaete-scute homolog 3 (bHLH transcriptional regulator Sgn-1)
		<u> </u>	Alternate: ASCL3
	NM_054048	U:2.16	RE1-silencing transcription factor (REST) co-repressor; co-repressor of Rest; Rest
	NP 473389.1	(YtoO)	co-repressor
	111_473369.1	(1100)	co-repressor
	NM_016968	U:2.16	
	NP_058664.1	(YtoO)	Oligodendrocyte transcription factor 1 (Oligo1).
	AF316872	U:2.16	
	AAK28061.1	(YtoM)	PTEN induced putative kinase 1; protein kinase BRPK
	1.605.5	77.6 1 5	
	M62766	U:2.16	·
	AAA37819.1	(YtoM)	3-hydroxy-3-methylglutaryl-coenzyme A reductase (HMG-CoA reductase).

	.1	1	1
	AK006525	U:2.16	
	BAB24634.1	(YtoM)	Islet cell autoantigen
			Subclass: islet cell autoantigen p69 (diabetes-associated autoantigen p69)
			Subclass: islet cell autoantigen 1 isoform 1; islet cell autoantigen 1 (69kD); islet ce
5			autoantigen p69
			Subclass: islet cell autoantigen 1 isoform 2; islet cell autoantigen 1 (69kD); islet ce
			autoantigen p69
	NM_009350	U:2.15	
	NP 033376.1	(MtoO)	Nuclear RNA-binding protein
0	033370.1	(Wilou)	Subclass: testis nuclear RNA-binding protein
_			Substituti 14 11 Olitalis protein
	NM_018779	U:2.15	phosphodiesterase
	NP_061249.1	(5to19)	·
			Subclass: cGMP-inhibited 3',5'-cyclic phosphodiesterase A (Cyclic GMP inhibited
	`		phosphodiesterase A) (CGI-PDE A)
5			Subclass: phosphodiesterase 3.A, cGMP-inhibited
		,	Subclass: cGMP-inhibited cAMP phosphodiesterase (EC 3.1.4), myocardial form
			Subclass: cyclic nucleotide phosphodiesterase
			Subclass: phosphodiesterase 3B, cGMP-inhibited
)	NM_011128	U:2.14	pancreatic lipase-related protein
	NP_035258.1	(5to11)	
			Subclass: pancreatic lipase-related protein 2
			Subclass: pancreatic lipase-related protein 1
			Alternate: pancreatic lipase
5			
	U36475	U:2.14	breast/ovarian cancer susceptibility protein BRCA1
	AAC52323.1	(5to11)	
	NM_007836	U:2.14	growth arrest and DNA-damage-inducible, alpha; DNA-damage-inducible transcrip
)	NP_031862.1	(5to19)	1; DNA damage-inducible transcript-1; DNA damage-inducible transcript 1
	NM_010361	U:2.14	glutathione S-transferase
	NP 034491.1	(5to19)	
	_	<u> </u>	Subclass: glutathione S-transferase theta 2
5		 	Subclass: glutathione S-transferase theta 1
,	`	 	Subclass: similar to Glutathione S-transferase theta 1 (GST class-theta) (Glutathion
			transferase T1-1)
		 	, ,

		1	4.1
	NM_008193	U:2.13	
	NP 032219.1	(YtoM)	Guanylate kinase (GMP kinase).
		1	Subclass: guanylate kinase 1
_	ND 4 013901	TT-2 12	
5	NM_013891	U:2.13	
	NP_038919.1	(MtoO)	prostate epithelium-specific Ets transcription factor
	NM_008542	U:2.13	MAD, mothers against decapentaplegic homolog 6; Mothers against
	NP 032568.1	(11to19)	decapentaplegic, drosophila, homolog of, 6; MAD (mothers against decapentaplegic,
	_032308.1	(11,015)	Drosophila) homolog 6
10		 	Alternate: MAD, mothers against decapentaplegic homolog 7; MAD (mothers
10		ĺ	against decapentaplegic, Drosophila) homolog 7; Mothers against decapentaplegic,
			drosophila, homolog of, 7
	NM_013607	U:2.13	mysosin heavy chain
	- NP 038635.1	(5to19)	
		(31017)	
		<u> </u>	Subclass: smooth muscle myosin heavy chain
15			Subclass: smooth muscle myosin heavy chain 11
		ļ	Subclass: smooth muscle myosin heavy chain 11, isoform SM1
			Subclass: smooth muscle myosin heavy chain 11, isoform SM2
		<u> </u>	Subclass: smooth muscle my osin heavy chain 11, isoform SM3
			Subclass: myosin, heavy polypeptide 9, non-muscle Subclass: Myosin heavy chain, nonmuscle type B (Cellular myosin heavy chain,
20			
			type B) (Nonmuscle myosin heavy chain-B) (NMMHC-B) Subclass: myosin heavy chain nonmuscle form A
		 	Subclass: nonmuscle myosin heavy chain (NMHC)
	·	 	Subclass: myosin, heavy polypeptide 7, cardiac muscle, beta
		 	Subclass: beta-myosin heavy chain
25		1	Subclass: myosin alpha heavy chain, cardiac muscle
2.0			Subclass: similar to cardiac a lpha-myosin heavy chain
			Subclass: Myosin heavy chain, fast skeletal muscle, embryonic (Muscle embryonic
			myosin heavy chain) (SMHCE)
			Subclass: cardiac beta myosin heavy chain
			Subclass: myosin heavy chaim, perinatal skeletal muscle
30			Subclass: similar to Myosin heavy chain, skeletal muscle, perinatal (MyHC-
			perinatal)
			Subclass: myosin, heavy poly peptide 1, skeletal muscle, adult; myosin heavy chain
			IIx/d
			Subclass: myosin, heavy polypeptide 2, skeletal muscle, adult
			Subclass: myosin, heavy polypeptide 3, skeletal muscle, embryonic
			Subclass: myosin, heavy polypeptide 4, skeletal muscle
35			Subclass: myosin, heavy polypeptide 8, skeletal muscle, perinatal

sophila,
sophila,
sophila,
sophila,
-
d circadian
T 1 7 C
; Inhibitor of
·
t negative
e S-
glutathione
: S-
T, class
ryltransferase
ase A2;
,
Ethacrynic
4;
glutathione
0
glutathione
111 121

·	1	412
NM_009075	U:2.09	· · · · · · · · · · · · · · · · · · ·
NP 033101.1	(YtoO)	Isomerase
		Subclass: ribose 5-phosphate isomerase A (ribose 5-phosphate epimerase);
		RIBOSE 5-PHOSPHATE ISOMERASE
NM_053082	U:2.09	
NP_444312.1	(YtoM)	Transmembrane protein
		Subclass: Transmembrane 4 superfamily, member 7 (Novel antigen 2) (NAG-2)
		(Tetraspanin 4) (Tspan-4).
		Subclass: tetraspan NET-5
NM_019670	U:2.09	
NP 062644.1	(MtoO)	Diaphanous protein homolog
NP_002044.1	(MIGO)	Subclass: Diaphanous protein homolog 3 (Diaphanous-related formin 3) (DRF3).
		Subclass: bA218B22.1.1 (novel protein (presumed ortholog of mouse diaphenous
		related formin (DIA2)) (isoform 1))
		Subclass: diaphanous 2 isoform 12C
		Subclass: diaphanous 2 isoform 156
		Subclass: Diaphanous protein homolog 2 (Diaphanous-related formin 2) (DRF2).
		Subclass: diaphanous 1; Diaphanous, Drosophila, homolog of, 1; deafness,
		autosomal dominant 1; diaphanous (Drosophila, homolog) 1; hDia1
		Subclass: bA218B22.1.2 (novel protein (presumed ortholog of mouse diaphenous
	1	related formin (DIA2)) (translation of cDNA DKFZp434C0931 (Em:AL137718))
	 	(isoform 2))
√M_011066	U:2.08	
_	(YtoO)	Circa dian areatain hamalas
NP_035196.1	(1100)	Circadian protein homolog Subclass: period 2 isoform 2; period, Drosophila, homolog of, 2; period circadian
		protein 2
	1	Subclass: period 2 isoform 1; PERIOD, DROSOPHILA, HOMOLOG OF, 2; period 2 isoform 1; PERIOD, DROSOPHILA, HOMOLOG OF, 2; period 2 isoform 1; PERIOD, DROSOPHILA, HOMOLOG OF, 2; period 2 isoform 1; PERIOD, DROSOPHILA, HOMOLOG OF, 2; period 2 isoform 1; PERIOD, DROSOPHILA, HOMOLOG OF, 2; period 2 isoform 1; PERIOD, DROSOPHILA, HOMOLOG OF, 2; period 2 isoform 1; PERIOD, DROSOPHILA, HOMOLOG OF, 2; period 2 isoform 1; PERIOD, DROSOPHILA, HOMOLOG OF, 2; period 2 isoform 1; PERIOD, DROSOPHILA, HOMOLOG OF, 2; period 2 isoform 1; PERIOD, DROSOPHILA, HOMOLOG OF, 2; period 2 isoform 1; PERIOD, DROSOPHILA, HOMOLOG OF, 2; period 2 isoform 1; PERIOD, DROSOPHILA, HOMOLOG OF, 2; period 2 isoform 1; PERIOD, DROSOPHILA, HOMOLOG OF, 2; period 2 isoform 1; PERIOD, DROSOPHILA, HOMOLOG OF, 2; period 2 isoform
		circadian protein 2
		Subclass: period 1; period (Drosophila) homolog 1; hPER; Period, drosophila,
		homolog of; circadian pacemaker protein RIGUI
		Subclass: Period circadian protein 3 (hPER3).
	<u> </u>	Subclass: KIAA0482 protein
AK017753	U:2.08	
CP 285418	(YtoO)	Zinc finger protein
AB041576	U:2.08	
	1	Hydrolase
3AA95060.1	(YtoO)	
	ı	Subclass: nudix (nucleoside diphosphate linked moiety X)-type motif 11;

			414
			Subclass: nudix (nucleoside diphosphate linked moiety X)-type motif 4
			Subclass: diphosphoinositol polyphosphate phosphohydrolase type 2 beta
			Subclass: diphosphoinositol polyphosphate phosphohydrolase type 2 alpha
5	S70056	U:2.08	
	AAB30620.2	(YtoM)	Reductase
		(=====,	Subclasscrystallin, zeta; quinone oxidoreductase; NADPH:quinone reductase
	-	 	Substitution of the substi
	L25890	U:2.08	
0	AAA72411.1	(YtoM)	Receptor protein-tyrosine kinase
0	MAN /2411.1	(TIOIVI)	Subclass: ephrin receptor EphB2 isoform 1 precursor; developmentally-regulated
			eph-related tyrosine kinase; elk-related tyrosine kinase; eph tyrosine kinase 3
			Subclass: Ephrin type-B receptor 2 precursor (Tyrosine-protein kinase receptor
	· ·	1	
		 	EPH-3) (DRT) (Receptor protein-tyrosine kinase HEK5)(ERK). Subclass: ephrin receptor EphB2 isoform 2 precursor; developmentally-regulated
	ļ	 	eph-related tyrosine kinase; elk-related tyrosine kinase; eph tyrosine kinase 3
		1	Subclass: ephrin receptor EphB1 precursor; eph tyrosine kinase 2; ephrin receptor
			EphB1
		ł	Subclass: ephrin receptor EphB3 precursor; human embryo kinase 2; EPH-like
5		ļ	tyrosine kinase 2; tyrosine-protein kinase receptor HEK-2
		<u> </u>	Subclass: Eph-like receptor tyrosine kinase hEphB1c
		<u> </u>	Subclass: EphA4; Hek8; TYRO1 protein tyrosine kinase; ephrin receptor EphA4
			Subclass: EphA7; Hek11; ephrin receptor EphA7
		1	Subclass: EphA5; Hek7; ephrin receptor EphA5
			Subclass: EphA3; Ephrin receptor EphA3 (human embryo kinase 1); eph-like
)			tyrosine kinase 1 (human embryo kinase 1); ephrin receptor EphA3
			Subclass: ephrin receptor EphB4 precursor; Ephrin receptor EphB4 (hepatoma
	1		transmembrane kinase); Tyro11; ephrin receptor EphB4; hepatoma transmembrane
			ikinase
			Subclass: ephrin receptor EphA8 precursor; ephrin type-A receptor 8precursor; eph-
			and elk-related tyrosine kinase; tyrosylprotein kinase; tyrosine-protein kinase
		1	
		 	receptor eek; protein-tyrosine kinase; hydroxyaryl-protein kinase Subclass: ephrin receptor EphB6 precursor; tyrosine-protein kinase-defective
		 	receptor; ephrin type-B receptor 6 EphA2; ephrin receptor EphA2; epithelial cell receptor protein tyrosine
			EphA2; ephrin receptor EphA2; epithelial cell receptor protein tyrosine
			kinase
			EphA1; eph tyrosine kinase 1 erythropoietin-producing hepatoma amplified
			sequence; oncogene EPH; ephrin receptor EphA1); ephtyrosine kinase 1
		<u> </u>	(erythropoietin-producing hepatoma amplified sequence); ephrin receptor EphA1
			Subclass: ephrin receptor EPHA3 secreted form
			Subclass: hemopoietic cell kinase
			Subclass: protein-tyrosine kinase hck
			Subclass: viral oncogene yes-1 homolog 1; proto-oncogene tyrosine-protein kinase
		1	YES; Yamaguchi sarcoma oncogene; cellular yes-1 protein
			Subclass: PTK2 protein tyrosine kinase 2 isoform a; focal adhesion kinase 1
			Subclass: PTK2 protein tyrosine kinase 2 isoform b; focal adhesion kinase 1
		<u> </u>	protein tylosino kiliaso r isotorino, todar adirestori kiliaso r

			415
			Subclass: protein-tyrosine kinase fyn isoform a; proto-oncogene tyrosine-protein
		1	kinase fyn; src/yes-related novel gene; src-like kinase; c-syn protooncogene; tyrosine
			kinase p59fyn(T);OKT3-induced calcium influx regulator
			Subclass: lymphocyte-specific protein tyrosine kinase; oncogene LCK; membrane
			associated protein tyrosine kinase
			Subclass: v-src sarcoma (Schmidt-Ruppin A-2) viral oncogene homolog;
			Protooncogene SRC, Rous sarcoma; v-src avian sarcoma(Schmidt-Ruppin A-2) viral
			oncogene hornolog
			Subclass: v-abl Abelson murine leukemia viral oncogene homolog 1 isoform b;
	<u></u>		Abelson murime leukemia viral (v-abl) oncogene homolog 1
	1.		Subclass: v-abl Abelson murine leukemia viral oncogene homolog 1 isoform a;
5			Abelson murime leukemia viral (v-abl) oncogene homolog 1
			Subclass: protein-tyrosine kinase fyn isoform b; proto-oncogene tyrosine-protein
			kinase fyn; src/yes-related novel gene; src-likekinase; c-syn protooncogene; tyrosine
			kinase p59fyn(T); OKT3-induced calcium influx regulator
	1		Subclass: protein-tyrosine kinase fyn isoform c; proto-oncogene tyrosine-protein
			kinase fyn; src/yes-related novel gene; src-likekinase; c-syn protooncogene; tyrosine
			kinase p59fyn(T);OKT3-induced calcium influx regulator
•			Subclass: fer (fps/fes related) tyrosine kinase (phosphoprotein NCP94); fer (fps/fes
			related) tyrosime kinase
		<u> </u>	Subclass: tec protein tyrosine kinase
10		-	Subclass: protein-tyrosine kinase (EC 2.7.1.112) FAK - human
•			Subclass: v-ab1 Abelson murine leukemia viral oncogene homolog 2 isoform a;
			Abelson-related protein; arg
			Subclass: v-ab1 Abelson murine leukemia viral oncogene homolog 2 isoform b; arg;
	·	 	Abelson murine leukemia viral (v-abl) oncogene homolog 2 (arg,
	77.602120	TX 2 00	
	NM_023128	U:2.08	
15	NP_075617.1	(MtoO)	Plasma membrane protein
			Subclass: Paralemmin
		+	Subclass: KIA A0270 Subclass: para lemmin 2
		 	Subclass: Palm2-AKAP2 fusion protein
20			Subclass. 1 and 2-ARAF 2 fusion protein
20	NM_030696	U:2.08	solute carrier family 16 (monocarboxylic acid transporters)
	NP_109621.1	(5to19)	
		(3.015)	
	•		Subclass: solute carrier family 16 (monocarboxylic acid transporters), member 1;
		 	Solute carrier family 16 (monocarboxylic acid transporters),
			Subclass: solute carrier family 16 (monocarboxylic acid transporters), member 3;
0 =			monocarboxylate transporter 3 Subclass: solute corrier family 16 (managerhoxylic acid transporters) manha 7.
25			Subclass: solute carrier family 16 (monocarboxylic acid transporters), member 7;
			monocarboxylate transporter 2 Subclass: monocarboxylate transporter 1
-			Substass. Inditocarboxytate transporter 1
	L		

			. 416
	NM_022888	U:2.08	
	NP 075026.1	(5to19)	folate receptor
	1	(3333)	Subclass: folate receptor 1 (adult)
			Subclass: folate receptor 3
5			Subclass: folate binding protein
	NM_019811	U:2.07	
	NP 062785.1	(YtoM)	Acetyl-CoA synthetase
	3	(/ -	Subclass: acetyl-CoA synthetase isoform a; cytoplasmic acetyl-coenzyme A
			synthetase; acetate-CoA ligase; acyl-activating enzyme; acetate thiokinase; acetyl-
			CoA synthetase
		1	Subclass: acetyl-CoA synthetase isoform b; cytoplasmic acetyl-coenzyme A
			synthetase; acetate-CoA ligase; acyl-activating enzyme; acetate thiokinase; acetyl-
0			CoA synthetase
			Subclass: dJ568C11.3 (novel AMP-binding enzyme similar to acetyl-coenzyme A
		<u> </u>	synthethase (acetate-coA ligase))
			Subclass: KIAA1846 protein
			Subclass: dJ18C9.1.3 (similar to acetyl-coenzyme A synthetase, isoform 3)
		 	
5	NM_016675	U:2.06	
	NP 057884.1	(YtoM)	Claudin
			Subclass: claudin 2
	NM_011414	U:2.05	·
0	NP 035544.1	(YtoO)	Protease inhibitor
			Subclass: secretory leukocyte protease inhibitor; antileukoproteinase; seminal
		-	proteinase inhibitor; mucus proteinase inhibitor
	U89924	U:2.05	protein phosphatase 1, regulatory (inhibitor) subunit 5; Phosphatase 1, regulatory
	AAB49689.1	(5to7)	inhibitor subunit 5
5			
	NM_016878	U:2.04	·
	NP 058574.1	(YtoM)	Aminopeptidase
_			Subclass: aspartyl aminopeptidase
)	NM_010421	U:2.04	
	NP 034551.1	(YtoM)	Aminidase
		1	Subclass: hexosaminidase A preproprotein; beta-hexosaminidase alpha chain; beta-
			N-acetylhexosaminidase: N-acetyl-beta-glucosaminidase
		<u> </u>	
5		 	
0	NM_010421 NP_034551.1	U:2.04 (YtoM)	Aminidase Subclass: hexosaminidase A preproprotein; beta-hexosaminidase alpha chain; beta-nexosaminidase; N-acetyl-beta-glucosaminidase Subclass: N-acetyl-alpha-glucosaminidase Subclass: N-acetyl-beta-glucosaminidase Subclass: hexosaminidase B preproprotein; beta-hexosaminidase beta chain; beta

		•	417
			acetylhexosaminidase; N-acetyl-beta-glucosaminidase
		<u> </u>	
	NM_019430	U:2.04	
•	NP 062303.1	(YtoM)	Valtaga danandant galajum ahannal
	NP 062303.1	(I tolvi)	Voltage-dependent calcium channel Subclass: voltage-dependent calcium channel gamma-3 subunit; neuronal voltage-
			gated calcium channel gamma-3 subunit Subclass: voltage-dependent calcium channel gamma-2 subunit; stargazin; neuronal
4.0			
40			voltage-gated calcium channel gamma-2 subunit Subclass: voltage-dependent calcium channel gamma-4 subunit; neuron al voltage-
			1
		<u> </u>	gated ca1cium channel gamma-4 subunit Subclass: voltage-dependent calcium channel gamma-8 subunit; neuron al voltage-
			1
			gated ca1cium channel gamma-8 subunit
			
	NM_010444	U:2.04	
45	NP 034574.1	(MtoO)	Nuclear receptor
			Subclass: nuclear receptor subfamily 4, group A, member 1 isoform a; hormone
			receptor; growth factor-inducible nuclear protein N10; early response protein NAK1;
			orphan nuclear receptor HMR; TR3 orphan receptor; steroid receptor TR3
			Subclass: TR3 orphan receptor
		-	Subclass: nuclear receptor subfamily 4, group A, member 2 isoform a; nur related
		1	protein-1 (mouse), human homolog of; transcriptionally inducible nuclear receptor
		Į	
		1	related 1; intermediate-early receptor protein; T-cell nuclear receptor NOT; orphan
			nuclear receptor NURR1; NGFI-B/nur77 beta-type transcription factor homolog
			Subclass: nuclear receptor subfamily 4, group A, member 2 isoform d; nur related
			protein-1 (mouse), human homolog of; transcriptionally inducible nuclear receptor
			related 1; intermediate-early receptor protein; T-cell nuclear receptor NOT; orphan
			nuclear receptor NURR1; NGFI-B/nur77 beta-type transcription factor homolog
50			Subclass: NGFI-B/nur77 beta-type transcription factor homolog
			Subclass: nuclear receptor subfamily 4, group A, member 2 isoform b; nur related
			protein-1 (mouse), human homolog of; transcriptionally inducible nuclear receptor
			related 1; intermediate-early receptor protein; T-cell nuclear receptor NOT; orphan
			nuclear receptor NURR1; NGFI-B/nur77 beta-type transcription factor homolog Subclass: Nuclear hormone receptor NOR-1 (Neuron-derived orphan receptor 1)
		 	(Mitogen induced nuclear orphan receptor). Subclass: nuclear receptor subfamily 4, group A, member 3 isoform b;
			chondros arcoma, extraskeletal myxoid, fused to EWS; translocated in extraskeletal
			chondrosarcoma; neuron derived orphan receptor; mitogen induced nuclear orphan
			receptor
			Subclass: steroid/thyroid orphan receptor homolog gene
			Subclass: nuclear receptor subfamily 4, group A, member 3 isoform a;
		1	chondros arcoma, extraskeletal myxoid, fused to EWS; translocated in extraskeletal
		1	chondros arcoma; neuron derived orphan receptor; mitogen induced nucle ar orphan
55			receptor
رر	<u> </u>	<u> </u>	**Coopioi

		_	418
			Subclass: mitogen induced nuclear orphan receptor
			Subclass: nuclear receptor subfamily 4, group A, member 1 isoform b; hormone
			receptor; growth factor-inducible nuclear protein N10; early response protein NAK1
			orphan nuclear receptor HMR; TR3 orphan receptor; steroid receptor TR3
			Subclass: nuclear receptor subfamily 4, group A, member 2 isoform c; mur related
	·		protein-1 (mouse), human homolog of; transcriptionally inducible nuclear receptor
			related 1; intermediate-early receptor protein; T-cell nuclear receptor NOT; orphan
			nuclear receptor NURR1; NGFI-B/nur77 beta-type transcription factor homolog
5	NM_023740	U:2.04	PP3774
	NP_076229.1	(5to11)	
			Alternate: Similar to RIKEN cDNA 1500015N03 gene
			Alternate: similar to Abl-philin 2
			Alternate: Abl-philin 2
10			
	NM_030566	U:2.04	Fos-related antigen
	NP_085043.1	(5to11)	
		U:2.04	Phosphormannomutase
	AK004631	1	Phosphorizannomidase
15	BAB23425.1	(5to19)	Subclass: Phosphomannomutase i (PMM 1) (PMMH-22)
		-	Subclass: phosphomannomutase 2
	NM 011498	U:2.03	
20	NP 035628.1	(YtoM)	Basic helix-loop-helix domain containing transcription factor
2.0	141 055028.1	(11011)	Subclass: differentiated embryo chondrocyte expressed gene 1
•			Subclass: bHLH transcription factor DEC1
			DECO.
			Subclass: basic helix-loop-helix domain containing, class B, 3; bHLH protein DEC2
25	NM_025703	U:2.03	
20	NP 079979.1	(YtoM)	hypothetical protein MGC45400
	111 075575.1	(1:01.1)	,
	AK012163	U:2.03	unnamed protein product
	BAB28070.1	(7to11)_	
30			Alternate: hypothetical protein FLJ10998
		 	
	NM_010368	U:2.02	
	NP 034498.1	(YtoM)	glucuroni dase
			Subclass: glucuronidase, beta
35			

	NM_009998	U:2.02	cytochrome P450, subfamily IIB (phenobarbital-inducible), polypeptide 6
	NP_034128.1	(11to19)	· ·
	ļ	+	
	NM_019692	U:2.02	Ras-Like without CAAX 2; Ric-like, expressed in neurons (Drosophila); GTP-
5	NP_062666.1	(5to19)	binding protein Roc2
		<u> </u>	Alternate: Ras family small GTP binding protein RIN
		 	Alternate: RIBA
			Alternate: Ras-like without CAAX 1; Ric-like, expressed in many tissues
-			(Dros ophila); GTP-binding protein Roc1
			(Brosopinia), GTI omaing protein recei
10	NM_025721	U:2.02	glycosylated 38 kDa sperm protein C-7/8 precursor
	NP_079997.1	(5to19)	
	NM_009708	U:2.01	
	NP 033838.1	(YtoO)	GTP-binding protein
15			Subclass: GTP-binding protein Rho7
			Subclass: ras homolog gene family, member E; Rho8; RhoE
		<u> </u>	Subclass: GTP-binding protein RHO6
٠.	NM_012042	U:2.01	·
20	NP_036172.1	(YtoM)	Cullin proteins
			Subclass: cullin 1
			Subclass: cullin 2
		-	Subclass: cullin 3
		,	Subclass: Vasopressin-activated calcium-mobilizing receptor (VACM-1) (Cullin
0.5	ļ		homolog 5) (CUL-5).
25			Subclass: cullin 4B; Cullin-4B
			Subclass: cullin 4A
	AK010827	U:2.01	
	BAB27209.1	(YtoM)	hypothetical protein FLJ12660
30		<u> </u>	
	NM_010107		ephrin A1; eph-related receptor tyrosine kinase ligand 1 (turnor necrosis factor,
	NP_034237.1	(5to7)	alpha-induced protein 4)
	NM_011710	U:2	
35	NP 035840.1		tRNA synthetase
		1	Subclass: Tryptophanyl-tRNA synthetase (TryptophantRNA ligase) (TRPRS);
		<u> </u>	interferon-induced protein 53 (IFP53) (hWRS).

NO TEXT

Mouse Gene	Behavior	Description
Protein		
AK004731	F:-2. O7 (YtoM)	
XP_148015	U:+2.71 (7to19)	plakophilin
		Subclass: plakophilin 2
		Subclass: plakophilin 2a
NM_009922	F:-2.54 (YtoO)	
NP_034052.1	U:+2.55 (7to19)	calponin
<u>.</u>		Subclass: calponin 1, basic, smooth muscle; calponins, basic; Calponin
		Subclass: calponin 2; Calonin 2
·		Subclass: calponin 3; calponin, acidic
NM_021291	F:-2. 1 1 (YtoM)	
NP_067266.1	U:+3.03 (5to19)	amino acid transporter
		Subclass: solute carrier family 7 (cationic amimo acid transporter, y+
		system), member 9; solute carrier family 7, member 9; solute carrier
		family 7 (cationic amino acid, transporter, y+ system), member 9
•		Subclass: solute carrier family 7 (cationic amino acid transporter, y+
		system), member 5; Membrane protein E16; So lute carrier family 7,
		member 5; 4F2 light chain
		Subclass: solute carrier family 7, (cationic amimo acid transporter, y+
		system) member 11; cystine/glutamate transporter
		Subclass: solute carrier family 7 (cationic amin o acid transporter, y+
		system), member 7
		Subclass: solute carrier family 7 (cationic amin o acid transporter, y+
		system), member 6
		Subclass: solute carrier family 7 (cationic amin o acid transporter, y+
		system), member 8
		Subclass: Y+L amino acid transporter 1 (y(+)L-type amino acid
		transporter 1) (y+LAT-1) (Y+LAT1) (Monocyte amino acid permease
		(MOP-2).
		Subclass: solute carrier family 7, member 10; a.sc-type amino acid
		transporter 1
		Subclass: Large neutral amino acids transporter small subunit 2 (L-typ
		amino acid transporter 2) (hLAT2).
NM_033373	F:-2.O5 (YtoO)	
NP 203537.1	U:+2.12 (7to19)	keratin

		422		
		Subclass: keratin 23 isoform a; hyperacetylation-inducible type I keratin;		
		keratin, type I cytoskeletal 23; cytokeratin 23; type I intermediate filament		
		cytokeratin; histone deacetylase inducible keratin 23, (Cytokeratin 23)		
		(K23) (CK 23).		
		Subclass: keratin 23 isoform b; hyperacetylation-inducible type I keratin;		
		keratin, type I cytoskeletal 23; cytokeratin 23; type I intermediate filament		
		cytokeratin; histone deacetylase inducible keratin 23		
		Subclass: keratin 20, type I-like, cytoskeletal		
		Subclass: keratin 19; keratin, type I cytoskeletal 19; keratin, type I, 40-kd;		
		cytokeratin 19; (Cytokeratin 19) (K19) (CK 19).		
	 	Subclass: keratin 17		
		Subclass: keratin 17 Subclass: keratin 12 (Meesmann corneal dystrophy); Keratin-12; keratin		
		12		
		Subclass: keratin 15; keratin-15, basic; keratin-1 5, beta; type I		
		cytoskeletal 15; cytokeratin 15; (Cytokeratin 15) (K15) (CK 15).		
	_	Subclass: keratin 13; keratin, type I cytoskeletal 13; cytokeratin 13		
		Subclass: keratin 16; keratin, type I cytoskeletal 16; cytokeratin 16		
		Subclass: keratin 14; cytokeratin 14		
		Subclass: type I hair keratin 6; keratin, hair, acidic, 6		
·		Subclass: cytokeratin 20		
		Subclass: type I hair keratin 5; Ha-5; hard keratin, type I, 5		
		Subclass: Keratin, type I cytoskeletal 10 (Cytokeratin 10) (K10) (CK 10).		
		Subclass: type I hair keratin 3A; Ha-3I; hard keratin, type I,3I; keratin,		
<u></u>		hair, acidic,3A		
		Subclass: type I hair keratin 1; hard keratin, type I, 1; Ha-1; keratin, hair,		
		acidic,1		
		Subclass: type I hair keratin 4; hard keratin, type I, 4		
		Subclass: type I hair keratin 2; Ha-2; hard keratin, type I, 2; keratin, hair,		
		acidic,2		
		Subclass: keratin 18		
		Subclass: cytokeratin 9		
X93035	F:-2.87 (YtoO)			
CAA63603.1	U:+2.78 (5to19)	chitinase		
		Subclass: chitinase 3-like 1; cartilage glycoprote in-39		
		Subclass: chitotriosidase; plasma methylumbelli feryl		
		tetra-N-acetylchitotetraoside hydrolase		
		Subclass: chitinase 3-like 2; chondrocyte proteira 39		
		Alternate: oviductal glycoprotein		
	1	Subclass: oviductal glycoprotein 1, 120kDa (mucin 9, oviductin); mucin		
		9 (oviductin); oviductal glycoprotein 1, 120kD (rmucin 9, oviductin)		
		(
	1			

1ŏ

NM_010444	U:2.04 (MtoO)			
NP_034574.1	F:-2.6 (7to11)	Nuclear receptor		
		Subclass: nuclear receptor subfamily 4, group A, member 1 isoform a;		
		hormone receptor; growth factor-inducible muclear protein N10; early		
		response protein NAK1; orphan nuclear receptor HMR; TR3 orphan		
		receptor; steroid receptor TR3		
		Subclass: TR3 orphan receptor		
		Subclass: nuclear receptor subfamily 4, group A, member 2 isoform a;		
		nur related protein-1 (mouse), human homolog of; transcriptionally		
		inducible nuclear receptor related 1; intermediate-early receptor protein		
		T-cell nuclear receptor NOT; orphan nuclear receptor NURR1;		
		NGFI-B/nur77 beta-type transcription factor homolog		
		Subclass: nuclear receptor subfamily 4, group A, member 2 isoform d;		
		nur related protein-1 (mouse), human homolog of; transcriptionally		
		inducible nuclear receptor related 1; intermediate-early receptor protein;		
		T-cell nuclear receptor NOT; orphan nuclear receptor NURR1;		
		NGFI-B/nur77 beta-type transcription factor homolog		
	1	Subclass: NGFI-B/nur77 beta-type transcription factor homolog		
		Subclass: nuclear receptor subfamily 4, group A, member 2 isoform b;		
		nur related protein-1 (mouse), human homolog of; transcriptionally		
	·	inducible nuclear receptor related 1; intermediate-early receptor protein		
		T-cell nuclear receptor NOT; orphan nuclear receptor NURR1;		
3		NGFI-B/nur77 beta-type transcription factor homolog		
		Subclass: Nuclear hormone receptor NOR-1 (Neuron-derived orphan		
	1	receptor 1) (Mitogen induced nuclear orphan receptor).		
		Subclass: nuclear receptor subfamily 4, group A, member 3 isoform b;		
		chondrosarcoma, extraskeletal myxoid, fused to EWS; translocated in		
		extraskeletal chondrosarcoma; neuron derived orphan receptor; mitogen		
		induced nuclear orphan receptor		
		Subclass: steroid/thyroid orphan receptor homolog gene		
		Subclass: nuclear receptor subfamily 4, group A, member 3 isoform a;		
		chondrosarcoma, extraskeletal myxoid, fused to EWS; translocated in		
		extraskeletal chondrosarcoma; neuron derived orphan receptor; mitogen		
		induced nuclear orphan receptor		
<u>, ., ., ., ., ., ., ., ., ., ., ., ., .,</u>	<u> </u>	Subclass: mitogen induced nuclear orphan receptor		
		Subclass: nuclear receptor subfamily 4, group A, member 1 isoform b;		
		hormone receptor; growth factor-inducible nuclear protein N10; early		
		response protein NAK1; orphan nuclear receptor HMR; TR3 orphan		
		receptor; steroid receptor TR3		

		424
		Subclass: nuclear receptor subfamily 4, group A, member 2 isoform c;
		nur related protein-1 (mouse), human homolog of; transcriptionally
		inducible nuclear receptor related 1; intermed iate-early receptor protein;
		T-cell nuclear receptor NOT; orphan nuclear receptor NURR1;
		NGFI-B/nur77 beta-type transcription factor homolog
NM_010831	U:3.91 (YtoO)	
NP 034961. 1	F:-2.39 (11to19)	Serine/threonine protein kinase
		Subclass: SNF1-like kinase
		Subclass: Ser/Thr protein kinase PAR-1A
·		Subclass: Ser/Thr protein kinase PAR-1B alpha
		Subclass: MAP/microtubule affinity-regulating kinase like 1; MARK4
		serine/threonine protein kinase
		Subclass: MAP/microtubule affinity-regulating kinase 2 isoform a; ELKI
		motif kinase 1; ELKL motif kinase
	·	Subclass: MAP/microtubule affinity-regulating kinase 2 isoform b;
		ELKL motif kinase 1; ELKL motif kinase
		Subclass: MAP/microtubule affinity-regulating kinase 3 long isoform
		Subclass: Cdc25C associated protein kinase C-TAK1
		Subclass: 5'-AMP-activated protein kinase, catalytic alpha-2 chain
		(AMPK alpha-2 chain).
		Subclass: KIAA0781 protein
		Subclass: KIAA0999 protein
NM_010846	U:2.39 (YtoO)	
NP_034976.1	F:-2.2 (5to7)	Interferon-induced protein
		Subclass: interferon-induced Mx protein
		Subclass: myxovirus resistance protein 1; interferon inducible protein
		p78; interferon-regulated resistance GTP-binding protein
		Subclass: interferon-induced viral resistance protein MxB
		Subclass: MX2
		Alternate: dynamin
		Subclass: dynamin 2; Dynamin II
		Subclass: Dynamin 3 (Dynamin, testicular) (T-dynamin).
NM_025703	U:2.03 (YtoM)	
NP_079979.1	F:-2.7 (5to19)	hypothetical protein MGC45400
ND 4 022174	TI-2 61 (V0)	
NM_033174	U:2.51 (YtoO)	DATE:
		1 A - 1 / 1 / 1 / 1
NP_149409.1	F:-2.07 (5to11)	snRNP Subclass: snRNP polypeptide B.

WO 2005/000335

PCT/US2004/017322

	,		
U67189	U:2.23 (YtoM)		
AAB50619.1	F:-3.57 (5to11)	regulator of G protein signalling 16	

15

20

25

Master Tables 101-199

In the related applications set forth at the beginning of the specification, we have looked at differential expression of genes in various organs and tissue with respect to (1) aging, (2) hyperinsulinemia and/or type II diabetes. Master Tables 101-199 (note that some of these table numbers are reserved for future use) tabulate those mouse genes which appear both in Master Table 1 of this application, and in the corresponding table of at least one of the related applications.

The following human proteins are considered to be of particular interest:

Human proteins corresponding to mouse genes listed as favorable both in Master Table 1 and in at least one of Master Tables 101-199, which are not listed as unfavorable in any of Master Tables 101-199; and

Human proteins corresponding to mouse genes listed as unfavorable both in Master Table 1 and in at least one of Master Tables 101-199, which are not listed as favorable in any of Master Tables 101-199.

WO 2005/000335 PCT/US2004/017322

•		427		
		Master Table 101	数据465570年上	de Salvingo.
*		MASCEL TABLE TO		
	Genes	Differentially Expressed With Respect:	to Age i	${f n}$.Both-
		Liver and Muscle		
			12,500	Muscle
	Mouse		Liver Aging	Aging
5	Gene	Mouse Description	Behavior	Behavior
_			A CONTRACTOR OF THE PARTY OF TH	
	77001045	Mus musculus 2-5A-dependent RNase L mRNA,	U:4.86	TT. 12 12
	AF281045	complete cds	(5tol1)	U:+2.12
	Ì			U:+2.26
		Mus musculus protein kinase BRPK mRNA,	U:2.16	
	AF316872	complete cds	(YtoM)	F:3.65
		AK015750 Mus musculus adult male testis	79-21	
	<u>}</u>	cDNA, RIKEN full-length enriched library,	į	Carrier Carrie
		clone:4930511F10:sulfotransferase, estrogen	U:2.56	
	AK015750	preferring, full insert sequence	(YtoO)	U:+7.39
)	Mus musculus adult male medulla oblongata		
	1	cDNA, RIKEN full-length enriched library,	U:4.01	
	AK018226	clone:6330533H24, full insert sequence	(5to19)	F:2.35
		MUSCOL1A4A Mus musculus alpha-1 type IV	F:2.05	!
10	J04694	collagen (Col4a-1) mRNA, complete cds	(5tol1)	F:6.66
		Mus musculus cell death-inducing DNA		
	NM 00770	fragmentation factor, alpha subunit-like	U:52.77	
	2	effector A (Cidea), mRNA	(YtoO)	U:+1.88
	NM 00795	TO THE RESIDENCE OF THE PROPERTY OF THE PROPER	F:2.65	
	2	kDa (Grp58), mRNA	(5to19)	F:2.59
			}	E.Z.J
15	· —	Mus musculus glutathione peroxidase 3	U:3.13	77 . 0 47
	1	(Gpx3), mRNA	(YtoO)	U:+2.43
	NM_00852		F:2.41	
	4	Mus musculus lumican (Lum), mRNA	(5to19)	F:2.01
	NM 00907	Mus musculus ribose 5-phosphate isomerase A	U:2.09	
20	5	(Rpia), mRNA	(YtoO)	F:2.48
	NM 00924	Mus musculus secreted acidic cysteine rich	F:2.73	
•	2 -	glycoprotein (Sparc), mRNA	(5to19)	F:4.66
	MM ODB38	Mus musculus thyroid hormone responsive	U:5.69	
•	1		K :	F:2.18
25	THE PROPERTY OF THE PARTY OF TH	THE PROPERTY OF THE PROPERTY O	STATE OF THE PARTY OF THE PARTY OF	PARTICULAR TRANSPORTATION OF THE PARTICULAR PROPERTY.
25		Mus musculus bromodomain-containing 2 (Brd2), mRNA	F:2.33 (7to19)	F:2.27
		(DLUZ) , IIIIVA middle Martine I i i i i i i i i i i i i i i i i i i	ر الاستون المعالمين مستحرين الاستثناء والمستدالية . أ	F . 4 . 4 /
	NM_01091	Annual Carl day	F:2.3	
	7	Mus musculus nidogen 1 (Nid1), mRNA	(5to11)	F:2.54
	i —		F:2.1	- 1
30	9	mrna	(5to19)	U:+2.72
	NM 01690	Mus musculus SEC61, alpha subunit (S.	F:2.37	U:+2.79
	6		(5to19)	F:3.89
	NM 01975	Mus musculus N-acetyltransferase 6 (Nat6),	F:2.02	
			(5to19) (F:2.55
35		4 A. C. L. L. L. L. L. L. L. L. L. L. L. L. L.		، سخانه مناهمه استراکی معادث میداد.
33	NM_01982	complex, subunit 3 (21 kDa) (Arpc3), mRNA	F:5.75	11. 42 74
	. 2012 1722 1	AS A THORN DELICATION OF THE CONTROL OF THE ASSESSMENT OF THE ASSE	~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~	
		Mus musculus solute carrier family 15	TT . 2 . 0.0	!
			F:3.08	E. 2 25
		CHARLES CONTRACTOR CON	(YtoM)	F:2.35
		Mus musculus cytochrome P450, subfamily IVF,		, i
4.0			F:2.19	TT . 0 . 7
40	4	hydroxylase) (Cyp4f14), mRNA	(5to19)	U:+2.12
	NM_02318	Mus musculus Kruppel-like factor 15 (Klf15),		U:+2.85
	4	mRNA	(5toll)	F:4.85

	Mus musculus RIKEN cDNA 2310005P05 gene (2310005P05Rik), mRNA	U:2.29 (5tol1)	U:+2.14
NM_02634 6	Mus musculus RIKEN cDNA 4833442G10 gene (4833442G10Rik), mRNA	F:3.64 (YtoO)	U:+6.12
	MMU89415 Mus musculus strain BALB/c elongation factor 2 mRNA, partial cds	F:2.73 (5to19)	

Table 102: Mouse Genes Differentially Expressed in Liver with respect to both Diabetes/Hyperinsulinemia and Aging

		Behavior	
Gene	Description	Diabetes	-Agi-ng
AF047725	Mus musculus CYP2C38 (Cyp2c38) mRNA, partial	F: (IR-D) 2.06 U: (C-D) 2.35	U:2.28 (5to l 1)
AF127033	Mus musculus fatty acid synthase mRNA, complete cds	F:(IR-D) 2.1	U:2.97 (YtoO)
AF294617	Mus musculus inducible 6-phosphofructo-2-kinase mRNA, complete cds	F:(C-IR) 2.63	F2.69 (5to 7)
AF385682	Mus musculus ETL1 mRNA, complete cds	F: (C-IR) 2.04, U: (IR-D) 2.02	F2.03 (7to 1 1)
Ar 30002	Mus musculus adult male kidney cDNA, RIKEN full-length enriched library,		
AK002693	clone:0610030A14:related to COSMID W01A11, full insert sequence	U: (C-IR) 2.04	(YtoO)
AK002979	Mus musculus adult male brain cDNA, RIKEN full-length enriched library, clone:0710001P07:homolog to D1 DOPAMINE RECEPTOR INTERACTING PROTEIN CALCYON, full insert sequence	F: (C-IR) 2.14, F: (C-D) 2.15	U:2. 6 7 (5to 1 9)
AK002979	Mus musculus adult male brain cDNA, RIKEN full-length enriched library, clone:07100C1P07:homolog to D1 DOPAMINE RECEPTOR INTERACTING PROTEIN CALCYON, full insert sequence	F: (C-IR) 2.14, F: (C-D) 2.15	U:2.67 (5to l 9)
AK005274	Mus musculus adult male cerebellum cDNA, RIKEN full-length enriched library, clone:1500017E18:homolog to HYDROXYACYLGLUTATHIONE HYDROLASE (EC 3.1.2.6) (GLYOXALASE II) (GLX II), full insert sequence	,	F3.89 (5to7)
AK005535	Mus musculus adult female placenta cDNA, RIKEN full-length enriched library, clone:1600025H15:homolog to CDNA FLJ20327 FIS, CLONE HEP10012, full insert sequence		F3.25 (YtoMI)
AK006096	AK006096 Mus musculus adult male testis cDNA, RIKEN full-length enriched library, clone:1700018018:hypothetical protein, full		U:4.75 (YtoO)
**************************************	clone:1700124F02:homolog to WUGSC:H_NH0335J18.1 PROTEIN, full insert	F: (C-IR) 2.95, U: (IR-D) 2.34	F2.04 (5tol 9)
AK007264	Mus musculus adult male testis cDNA, RIKEN full-length enriched library, clone:1700126L06:unclassifiable, full insert	U: (C-D) 2.19, U: (IR-D)	U:3.56
AK007293	Sequence Mus musculus adult male tongue cDNA, RIKEN	2.62 F:(C-IR)	(5tol 1)
AK009563	full-length enriched library, clone:2310032D16, full insert sequence		rz.⊥ (5tol 9)

5

.15

	430		
AK018226	Mus musculus adult male medulla oblongata cDNA, RIKEN full-length enriched library, clone:6330533H24, full insert sequence	F:(C-IR) 2.53, F:(C-D) 2.4	U:4.01
M12571	MUSHSP68A Mouse heat shock protein (hsp68) mRNA, clone MHS243, partial cds	U:(C-IR) 3.58	F2.73 (YtoM)
M12573	MUSHSP68C Mouse heat shock protein (hsp68) mRNA, clone MHS214, partial cds	U:(C-D) 2.94	F2.07 (5to19)
M62766	MUSHMGCOA Mouse HMG-CoA reductase mRNA, 3' end	U:(C-IR) 2.02	U:2.16 (YtoM)
M63245	MUSALASH Mus musculus amino levulinate synthase (ALAS-H) mRNA, 3' end	U:(C-IR) 3.05	F3.98 (5to19)
	Mus musculus apolipoprotein A-IV (Apoa4),	U (C-IR) 2.98, U (C-D) 2.42, U (IR-D)	F2 . 22
NM_007468	MRNA	2.16 F:(C-IR) 2.17,	(7to11)
NM_007472	Mus musculus aquaporin 1 (Aqp1), mRNA	U: (IR-D) 2.38	F2.04 (7to11)
NM_007489	Mus musculus aryl hydrocarbon receptor nuclear translocator-like (Arntl), mRNA	F:(C-D)- 2.13	F2.22 (7to11)
W. 007643	Mus musquing CD26 antigon (Cd26) mDNA	*	U: 3.57
January Commission of the Comm	Mus musculus CD36 antigen (Cd36), mRNA Mus musculus cell death-inducing DNA	3.33	(Yto0)
i i	fragmentation factor, alpha subunit-like effector A (Cidea), mRNA	U:(C-D)+ 4.7	U: 52.77 (YtoO)
	Mus musculus cytokine inducible SH2-containing protein 2 (Cish2), mRNA	F:(C-D) 2.51	F4 .4 (YtoM)
	Mus musculus carnitine acetyltransferase (Crat), mRNA	11 ' (U: 2.41 (5 to7)
NM_007809	Mus musculus cytochrome P450, 17 (Cyp17), mRNA	U: (C-IR) 3.41, U: (C-D) 3.69	U: 3.27 (YtoO)
	Mus musculus cytochrome P450, 26, retinoic acid (Cyp26), mRNA		F2 .08 (5 toll)
	Mus musculus cytochrome P450, 4a14	U: (C-IR) 24.5, F: (C-D) 5.06, F: (IR-D)	U: 18.8 (5 t 07)
NM 007824	Mus musculus cytochrome P450, 7al (Cyp7al),		U:2.47 (YtoM)

	431		
	Mara magallar subschape P450 7h1 (Cro7h1)	F: (C-IR) 6.41,	
NM_007825	Mus musculus cytochrome P450, 7bl (Cyp7bl), mRNA	U: (IR-D) 5.83 U: (C-IR)	(5to19)
	Mus musculus deiodinase, iodothyronine, type I (Dio1), mRNA	2.84,	F2.06 (7to19)
	Mus musculus epidermal growth factor receptor (Egfr), mRNA	F: (C-IR) 2.09, F: (C-D) 2.69	F2.21 (5to19)
	Mus musculus formyl peptide receptor, related sequence 2 (Fpr-rs2), mRNA	F:(C-D)-	The statement of the st
	Mus musculus glucose-6-phosphatase, catalytic (G6pc), mRNA		F2.75 (5toll)
	Mus musculus glutathione S-transferase, alpha 2 (Yc2) (Gsta2), mRNA	F: (C-IR) 9.17, F: (C-D) 5.68	U:5.76
NM_008245	Mus musculus hematopoietically expressed homeobox (Hhex), mRNA	F:(C-D) 2.62, U:(IR-D) 2.05	F2.2 (7to19)
	Mus musculus hydroxysteroid dehydrogenase-5, delta<5>-3-beta (Hsd3b5), mRNA	1	F2.25 (Yto0)
	Mus musculus insulin-like growth factor binding protein 1 (Igfbp1), mRNA	F:(C-IR) 3.37, F:(C-D) 3.47, F:(IR-D) 2.63	F13.28 (5to11)
	Mus musculus interleukin l beta (Illb), mRNA	F:(C-IR) 2.65, F:(C-D)	U = 3.05 (5to7)
	THE THE PARTY OF T	U:(C-IR) 2.59, F:(IR-D)	SALES SALES NEWS AND SALES OF
NM 008495	Mus musculus lectin, galactose binding, soluble 1 (Lgals1), mRNA	بأويسه والمساد والمسادات والمعادلة وخعاد	U: 4.6 (7toll)
at a Talente succession and a second	Mus musculus lipoprotein lipase (Lpl), mRNA	F: (C-D) 2.05, F: (IR-D) 2.42 U: (C-D)+	(5 to 19)
		2.68	(YtoO)

	432		
NM 009127	Mus musculus stearoyl-Coenzyme A desaturase 1 (Scd1), mRNA	F: (C-IR) 2.15, F: (C-D) 3.29, F: (IR-D) 2.71	U:2.2 (YtoM)
	Mus musculus serine protease inhibitor 4 (Spi4), mRNA	U: (IR-D) 2.01 F: (C-D) 2.61	U:3.6 (5to19)
Control of the second of the s	Mus musculus secreted phosphoprotein 1 (Spp1), mRNA	F:(C-IR) 2.04	F2.82 (5to19)
NM_009344	Mus musculus T-cell death associated gene (Tdag), mRNA	U: (IR-D) 2.1 F: (C-D) 3.91	F3.29 (7to19)
NM_009345	Mus musculus deoxynucleotidyltransferase, terminal (Dntt), mRNA	U: (C-D)+	U:2.43 (YtoO)
NM 009669	Mus musculus amylase 2, pancreatic (Amy2), mRNA	F: (C-IR) 3.13 U: (C-D) 3.23	F8.34 (5to7)
NM_009676	Mus musculus aldehyde oxidase 1 (Aox1), mRNA	-	U:2.36 (5to7)
	Mus musculus B-cell leukemia/lymphoma 6 (Bcl6), mRNA	2.11	F2.93 (5to19)
NM_009864	Mus musculus cadherin 1 (Cdh1), mRNA	*	F3.24 (YtoO)
NM 009895	Mus musculus cytokine inducible SH2-containing protein (Cish), mRNA	U: (IR-D) 2.45 F: (C-D) 2.25	F2.13 (Min)
	Mus musculus cytochrome P450, 2b10, phenobarbitol inducible, type b (Cyp2b10), mRNA	F: (C-IR) 2.61, F: (C-D) 2.33	U:2.02 (11to19)
	Mus musculus decay accelerating factor l (Daf1), mRNA	F: (C-IR) 2.04, U: (IR-D) 2.14	F2.11 (7to11)
	Mus musculus deoxyribonuclease II alpha (Dnase2a), mRNA		U:2.89 (5toll)
NM_010107		F:(C-D) 2.18	i
	MusMusculus Fc receptor, IgG, low affinity	F: (C-IR) 2.18, U: (IR-D) 2.55	F2.28 (7to19)
NM_010225	Mus musculus forkhead box F2 (Foxf2), mRNA	ستنبي مناه السطاء ومرابع مما شفاء دامل ١٩٠٠ ومرابعة	U:2.42 (5toll)
	Mus musculus glucocorticoid-induced leucine zipper (Gilz), mRNA		F3.32 (5to19)

10

10

	433		
NM_010324	Mus musculus glutamate oxaloacetate transaminase 1, soluble (Got1), mRNA	F: (C-D) 2.01	F2.08 (5tol1)
NM_010354	Mus musculus gelsolin (Gsn), mRNA	U: (C-IR) 2.03	F2.34 (5to19)
NM_010357	Mus musculus glutathione S-transferase, alpha 4 (Gsta4), mRNA	F: (C-IR) 2.17, F: (C-D) 2.93	U:2.11 (5to19)
NM_010361	Mus musculus glutathione S-transferase, theta 2 (Gstt2), mRNA	F: (C-IR) 2.46, F: (C-D) 2.25	U:2.14 (5to19)
NM_010634	Mus musculus fatty acid binding protein 5, epidermal (Fabp5), mRNA	U: (C-IR) 3.17, F: (IR-D) 5.62	F2.84 (5to19)
NM_011087	Mus musculus paired-Ig-like receptor Al (Piral), mRNA	2.49	F2.03 (YtoO)
NM_011125	Mus musculus phospholipid transfer protein (Pltp), mRNA	F: (C-IR) 2.01	U:3.1 (YtoO)
	Mus musculus pancreatic lipase-related	U: (C-D) 2.35, U: (IR-D) 2.73 F: (C-D)	U:2.14
NM_011128	protein 2 (Pnliprp2), mRNA	2.85	(5tol1) U:2.68
NM_011146	Mus mus culus peroxisome proliferator activated receptor gamma (Pparg), mRNA	F:(C-IR) 2.17	(5tol1)
NM_011375	Mus musculus sialyltransferase 9 (CMP-NeuAc:lactosylceramide alpha-2,3-sialyltransferase) (Siat9), mRNA	U:(C-IR) 2.65, U:(C-D) 2.16	F2.12 (5to19)
NM_011579	Mus musculus T-cell specific GTPase (Tgtp), mRNA	U:(C-IR) 2.13 F:(C-D) 2.1	F2.1 (5to19)
NM_011704	Mus musculus vanin 1 (Vnn1), mRNA	li '	p D
NM_012006	Mus musculus cytosolic acyl-CoA thioesterase 1 (Ctel), mRNA	F:(C-D) 2.24	U:3.07 (5to7)
NM_013459	Mus musculus adipsin (Adn), mRNA	F:(C-IR) 2.94	(5to11)
NM_013584	Mus musculus leukemia inhibitory factor receptor (Lifr), mRNA	والمتعلقين بدرسارات استكثار سيسدر	F3.35 (5to19)
NM_013594	Mus musculus methyl-CpG binding domain protein 1 (Mbd1), mRNA	U: (C-IR) 2.01, U: (C-D) 2.15	F2.35 (5to19)
NM_013623	Mus musculus orosomucoid 3 (Orm3), mRNA	U: (C-D)+	(7to19)
NM_013786		'U: (C-D)+	

	434		
		F: (C-IR) 3.7, U: (C-D)	F4.93
NM_015763	Mus musculus lipin 1 (Lpin1), mRNA	3.14	(5to19)
NM_016704	Mus musculus complement component 6 (C6),	F: (C-IR) 2.26, U: (IR-D) 3.29	F2.2 (5to19)
NM_016847	Mus musculus arginine vasopressin receptor 1A (Avprla), mRNA	U: (C-IR) 2.02, F: (IR-D) 2.03	F2.48 (5to19)
NM_016875	Mus musculus Y box protein 2 (Ybx2), mRNA	U: (IR-D) 2.73 F: (C-D) 4.72	F2.26 (YtoO)
NM_018779	Mus musculus phosphodiesterase 3A, cGMP inhibited (Pde3a), mRNA	F: (C-IR) 2.35, F: (C-D) 2.43	U:2.15 (5to19)
NM_018861	Mus musculus solute carrier family 1 (glutamate/neutral amino acid transporter), member 4 (Slc1a4), mRNA	U: (C-IR) 2.18	U:2.25 (YtoM)
NM_018887	Mus musculus cytochrome P450, 39a1 (Oxysterol 7alpha-hydroxylase) (Cyp39al-pending), mRNA	U:(C-D)+ 2.54	(7to19)
NM_019415	Mus musculus solute carrier family 12, member 3 (Slc12a3), mRNA	U:(C-IR) 2.06	U:2.6 (5to11)
NM_019811	Mus musculus acetyl-Coenzyme A synthetase 1 (AMP forming) (Acas1), mRNA	F: (C-IR) 2.03, F: (C-D) 2.11	U:2.07 (YtoM)
	Mus musculus cartilage associated protein (Crtap), mRNA	U: (C-D) 2.05 F: (C-D) 2.29	F2.03 (11to19)
NM_019977	-	U: (C-IR) 2.51 F: (C-D) 2.15	U:2.18 (YtoO)
		U: (C-IR) 2.06, U: (C-D) 2.23, U: (IR-D)	11.2 47
NM_019992	(Brdgl-pending), mRNA	2.12 U:(C-D)	(YtoO)
:	·	4.69	U:3.35 (5toll)
	Mus musculus sulfotransferase-related	F: (C-IR) 2.84, F: (C-D) 2.36, U: (IR-D)	
-: 	Mus musculus plasma membrane associated	U: (C-D)+ 2.12	(5to19) U:6.5 (YtoO)
	and the state of t		

10

Mus musculus unc13 homolog (C. elegans) 1 (Unc13h1), mRNA	F:(C-D)- 2.18	U:3.58 (MtoO)
Mus musculus homocysteine-inducible, endoplasmic reticulum stress-inducible, ubiquitin-like domain member 1 (Herpud1), mRNA	U: (C-IR) 3.00, U: (C-D) 2.29	F3.44 (5to19)
Mus musculus Kruppel-like factor 15 (Klf15), mRNA	U: (C-IR) 2.34	F2.87 (5to11)
Mus musculus camello-like 4 (Cml4), mRNA	F: (C-IR) 2.39, F: (C-D) 2.04	U:2.75 (5to19)
Mus musculus RIKEN cDNA 1500015N03 gene (1500015N03Rik), mRNA	F: (C-IR) 1.7, F: (C-D) 2.35, U: (IR-D) 2.52	U:2.04 (5tol1)
Mus musculus RIKEN cDNA 1110036H21 gene (1110036H21Rik), mRNA	F: (C-IR) 2.24, F: (C-D) 2.03	F3.11 (5tol1)
Mus musculus serine (or cysteine) proteinase inhibitor, clade B (ovalbumin), member 1a (Serpinbla), mRNA	F: (C-IR) 3.51, F: (C-D) 3.01	U:4.44 (5to19)
Mus musculus RIKEN cDNA 1700095F04 gene (1700095F04Rik), mRNA	F:(C-IR) 2.22	F2.72 (5to7)
Mus musculus RIKEN cDNA 2210418010 gene (2210418010Rik), mRNA	F:(C-D) 2.4	F2:28 (5to19)
Mus musculus type I intermediate filament cytokeratin (Haikl-pending), mRNA	4	F2.05 (YtoO)
Mus musculus RIKEN cDNA 0610033E06 gene (0610033E06Rik), mRNA	F: (C-IR) 1.98, F: (C-D) 3.23	F2.18 (5to19)
MMU67189 Mus musculus G protein signaling regulator RGS16 (rgs16) mRNA, complete cds	1 1	U:2.23 (YtoM)
MMU70139 Mus musculus probable nocturnin protein mRNA, partial cds	1	F2.05 (5to7)
		U:2.61 (YtoM)
	Mus musculus homocysteine-inducible, endoplasmic reticulum stress-inducible, ubiquitin-like domain member 1 (Herpudl), mRNA Mus musculus Kruppel-like factor 15 (Klf15), mRNA Mus musculus Kruppel-like factor 15 (Klf15), mRNA Mus musculus camello-like 4 (Cml4), mRNA Mus musculus RIKEN cDNA 1500015N03 gene (1500015N03Rik), mRNA Mus musculus RIKEN cDNA 1110036H21 gene (1110036H21Rik), mRNA Mus musculus serine (or cysteine) proteinase inhibitor, clade B (ovalbumin), member 1a (Serpinbla), mRNA Mus musculus RIKEN cDNA 1700095F04 gene (1700095F04Rik), mRNA Mus musculus RIKEN cDNA 2210418010 gene (2210418010Rik), mRNA Mus musculus type I intermediate filament cytokeratin (Haik1-pending), mRNA Mus musculus RIKEN cDNA 0610033E06 gene (0610033E06Rik), mRNA Mus musculus RIKEN cDNA 0610033E06 gene (0610033E06Rik), mRNA Mus musculus RIKEN cDNA 0610033E06 gene (0610033E06Rik), mRNA Mus musculus RIKEN cDNA 0610033E06 gene (0610033E06Rik), mRNA MMU67189 Mus musculus G protein signaling regulator RGS16 (rgs16) mRNA, complete cds MMU70139 Mus musculus probable nocturnin protein mRNA, partial cds MMU70139 Mus musculus probable nocturnin protein mRNA, partial cds	Mus musculus homocysteine-inducible, endoplasmic reticulum stress-inducible, ubiquitin-like domain member 1 (Herpudl), mRNA

10

5,

Table 201
Pairwise Differential Expression Comparisons

Selected Mouse Genes

5
_

10

Gene 🕹 🖠	Дде 5: 7.	Age 5 11	Age 35_19	:Age , 7-14-	Age 7-719	.Age 11,19	Age Y M	Äge Y 0	Age M 0
AK002979	U1.63	υ2.31	U2.94	U1.42	U1.81	U1.27	U2.90	U2.36	F1.23
AK004387	F1.79	F2.93	F3.29	F1.64	Fl.84	F1.12	F1.40	F2.33	F1.67
							U16.0	σ57.0	
NM_007702	U1.22	F1.07	U2.59	U1.30	U2.13	U2.78	9	1	U3.54
U67189	F2.04	F3.57	F1.91	F1.75	U3L.07	U1.86	F2.25	F1.02	υ2.21

Differential expression is set forth as the ratio of greater expression level to lesser expression level for the indicated time points. The direction of the change of expression is indicated by "F" (favorable, i.e., younger>older) or "U" (unfavorable, i.e., older>younger). Significant differences (at least two fold) are bold faced.

20

Note that in identifying a mouse gene as favorable, unfavorable, or mixed, only the significant (at least two fold) differentials are considered.

25 For the first six comparisons, the time points are weeks, e.g., "7 19" is 7 weeks vs. 19 weeks.

For the last three comparisons, the "Y", "M" and "O" represent

- 30 Y (young) = expression at 118 days
 - M (medium) = average of expression at 207 and 403 days
 - .0 (old) = average of expression at 558 and 725 daus

Example 2

5

10

15

20

25

30

35

The Amersham CodeLink™ Uniset Mouse I Bioarray Platform was used (example 1) to identify differences in liver gene expression in aging mice. The mice were fed normal chow and were sacrificed at ages ranging from 35 to 725 days. A total of 190 genes were differentially expressed by at least a 2-fold magnitude (Master Table 1). Analysis of the differentially expressed genes identified CIDE-A as the most differentially expressed gene in liver during this age span. The level of mouse CIDE-A expression in these mice is shown in figure 1.

437

No CIDE-A expression was detected at 35 to 56 days of age (expression level less than 0.2). The expression of CIDE-A was barely detectable at 118 and 207 days of age (0.36 +/- 0.23 and 0.23 +/- 0.10, respectively). However, CIDE-A is readily detected at 403 days of age (3.5 +/- 1.99) and the level of expression continues to increase to 7.7 (+/- 0.12) at 558 days of age. Taken together, the level of CIDE-A expression in liver increases at least 38-fold as the mouse progresses from 35 days of age to maximal expression at 558 days of age (7.7 +/- 0.12). See Figure 1.

The differentially expressed gene CIDE-A was subjected to further analysis.

Northern analysis

Total RNA (10 ug) from the appropriate tissues was resolved by denaturing agarose gel electrophoresis, transferred to positively charged nylon membrane, hybridized with the [α - 32 P]dCTP-labeled mouse CIDE-A cDNA (Random Primed DNA Labeling Kit, Roche, Indianapolis, IN) and exposed to Bio-Max MR film (Eastman Kodak Co., Rochester, NY).

Immunoblot analysis

Liver and heart tissue (100mg) was homogenized in 0.5ml phosphate buffered saline containing 7.5ul protease inhibitor cocktail (Sigma #P8340, St. Louis, MO). The samples were centrifuged for 5 min at 10,000 x g. The supernatant was

438

collected and protein concentration determined (Bio-Rad Laboratories #500-0006, Hercules, CA). Sixty micrograms of each extract was electrophoresed on a 12.5% SDS-polyacrylamide gel as described previously (25 Bowen). The resolved proteins were transferred to a nitrocellulose membrane and immunoblotted using a rabbit anti-mouse CIDE-A polyclonal antibody (QED Bioscience Inc., San Diego, CA) as previously described, see Kelder, B., Richmond, C., Stavnezer, E., List, E.O. and Kopchick, J.J., "Production, characterization and functional activities of v-Ski in cultured cells," Gene, 202:1521 (1997), and a goat anti-rabbit IgG polyclonal antibody conjugated to horseradish peroxidase.

15 Liver Histology

5

10

20

25

30

Liver tissues fixed in 4% paraformaldehyde were embedded in Tissue Path (Fisher Scientific, Pittsburgh, PA). Representative sections were prepared from each liver block, placed on a slide, subjected to H&E staining and evaluated by light microscopy. The percent white space was determined as a quantification of the level of steatosis.

Liver steatosis is observed in the CIDE-A expressing older mice.

We performed histological examinations on H&E stained liver sections prepared from mice of various ages to determine if increased CIDE-A expression effected any noticeable changes in the livers of these mice. Among other changes, we noticed an increased level of lipid accumulation within hepatocytes at 725 days of age. There was also an increased level of steatosis in liver tissue isolated from 558 day-old mice but the level of lipid accumulation did not approach that seen at 725 days.

35 CIDE-A is expressed at an early age in liver of high-fat fed type-II diabetic mice exhibiting liver steatosis.

Due to the correlation of increased CIDE-A expression and liver steatosis with increasing age, we investigated whether CIDE-A expression would also be increased in other models of liver steatosis. We utilized a mouse model of diet-induced obesity, hyperinsulinemia and type-II diabetes, see Surwit, R.S., Kuhn, C.M., Cochrane, C., McCubbin, J.A., Feinglos, M.N. (1988) "Diet-induced type-II diabetes in C57BL/6J mice," Diabetes 37:1163-1167. Mice were weaned onto either a normal diet or a high-fat diet for up to 26 weeks. Representative mice were sacrificed after 2, 4, 8, 16 and 26 weeks on the diet (35, 49, 77, 133 and 203 days of age) and CIDE-A expression levels were determined by DNA microarray analysis (Fig. 2).

We performed histological examinations on H&E stained liver sections prepared from control and type-II diabetic mice after 2, 16 and 26 weeks of high fat diet feeding (diet started at 3 weeks of age) to assess the degree of diet-induced liver steatosis (Fig. 3). The percent white space of each liver sample was determined by a histomorphometric profiling method using machine vision. H&E stained liver sections isolated from mice fed a normal diet at 56, 558 and 725 days of age shows the accumulation of lipid in liver hepatocytes of older mice.

Histological analysis indicated that diabetic liver hepatocytes accumulate a small amount of lipid as soon as 2 weeks on a high-fat diet and by 8 weeks, liver tissue isolated from high fat-fed mice contain significantly more lipid than their control counterparts. Severe liver steatosis is observed in liver tissues isolated from mice fed the high-fat diet for 16 weeks and is even more pronounced after 26 weeks of high-fat feeding. The percent white space in these livers is 31.6 and 53.2%, respectively. In comparison, the percent white space in liver tissue of mice fed the normal diet for 16 and 26 weeks is 10.3 and 12.2%, respectively. In addition, liver tissue isolated from 16 week high-fat fed hyperinsulinemic mice demonstrate liver steatosis but at a much lower level compared to its diabetic counterpart.

440

Correlation of CIDE-A gene expression and cell protein levels.

Since mRNA levels may not be indicative of the actual level of protein found in the tissue, we performed immunoblot analysis on heart and liver tissue isolated from control, hyperinsulinemic and type-II diabetic mice to confirm the increased CIDE-A levels.

Expression of genes involved in caspase-dependent apoptosis

Several groups have reported increase gene expression of members of the Caspase-dependent apoptotic pathway such as the FAS death receptor and Fas ligand in hepatocyte steatosis. See Feldstein, supra; Canbay A, Feldstein AE, Higuchi H, Werneburg N, Grambihler A, Bronk SF, Gores GJ. (2003) Kupffer cell engulfment of apoptotic bodies stimulates death ligand and cytokine expression. Hepatology 38:1188-1198. We therefore examined the levels of expression of genes involved in this pathway by DNA microarray analysis. A summary of the expression for the genes represented on the microarray is presented in Table 201.

Caspase-3 and -7

5

10

15

20

25

30

35

Expression levels of Caspase 3 and 7 both decrease from control to hyperinsulinemic to type-II diabetic. But immunohistochemistry on NASH liver sections and a rabbit antibody that recognizes a "neoepitope" (new epitope that is generated upon caspase 3 and 7 cleavage and activation) demonstrated increases in Caspase 3 and 7 activation. The decrease in caspase 3 and 7 gene expression may be an attempt by the cell to reduce apoptotic signaling within the cell (negative feedback).

Apoptosis in Liver

The level of apoptosis in liver may appear mirror. However the rapid phagocytosis of apoptotic bodies makes the detection of such bodies in tissue extremely difficult, see Savill, J. (2000) Apoptosis in resolution of inflammation. Kidney Blood

441

Press. Res. 23:173-174. A 4% rate of apoptosis would lead to a 25% reduction in liver tissue in 72 hours, see Schulte-Hermann, R., Bursch, W., Grasl-Kraupp, B. (1995) Active cell death (apoptosis) in liver biology and disease. Prog. Liver Dis. 13:1-35. Therefore, while it may be possible to observe only a small proportion of the ongoing apoptosis, the ongoing cell death may lead to major liver dysfunction.

Alternative Model

5

10

15

20

25

30

While increased apoptosis may be a contributing factor to liver dysfunction, we would like to put forth an alternate model for CIDE-A function in liver. In this model: CIDE-A is a part of a redundant apoptotic pathway. According to this model, in the early time points of the genesis of insulin resistance and Type-II diabetes, the liver is capable of managing liver steatosis by the primary caspase-activated apoptotic pathway to eliminate unwanted (lipid accumulating) However, as the disease progresses (and lipid hepatocytes. accumulates), the primary apoptotic pathway becomes overwhelmed (or non-functional) and a secondary (CIDE-A based) pathway is employed as an emergency (last-ditched) effort to maintain liver homeostasis However, this secondary, redundant apoptotic pathway that includes CIDE-A, is either not as efficient or incapable of eliminating the overwhelming lipid accumulation and eventual pathogenesis results.

It is possible that the apoptosis-induced cell death of lipid-containing hepatocytes results in the release of intracellular lipid and the concurrent extracellular liver lipid accumulation. This accumulation may then affect liver functions.

Raw expression values are stated, those resulting in 2-folg ofr greater differential expression The Amersham CodeLink $^{\text{TM}}$ high-fat fed mice exhibiting hyperinsulinemia or type-II diabetes after 16 week of feeding (N=2). Uniset Mouse I Bioarray Platform was used to determine the expression levels of control mice or Expression of genes involved in caspase-dependent apoptosis. are boldfaced. Table 201.

Type-II Diabetic	0.14 +/- 0.01	3.31 +/- 0.30	1.61 +/- 0.08	1.09 +/- 0.11	2.27 +/- 0.12	0.55 +/- 0.14	1.24 +/- 0.06	1.45 +/- 0.43	18.00 +/- 1.96	2.89 +/- 0.14	3.37 +/- 0.63	0.56 +/~ 0.03	54.59 +/- 4.64	0.20 +/- 0.03	1.76 +/- 0.19	0.46 +/~ 0.01	1.38 +/- 0.07	3.48 +/- 0.15	0.45 +/- 0.05	0.21 +/- 0.03	0.15 +/- 0.02
linemic	+/- 0.01	+/- 0.08	+/- 0.13	+/- 0.05	+/- 0.43	90.0 -/+	60-0 -/+	+/- 0.29	+/- 2.43	99.0 -/+	+/- 0.10	+/- 0.02	+/- 6.79	+/- 0.00	+/- 0.13	4/- 0.07	4/- 0.08	90.0 -/+	4/- 0.08	+/- 0.01	+/- 0.01
Hyperinsul	0.23	2.54	2.08	1.03	3.00	0,93	1.45	1.33	17.41	3.43	3.44	0.73	53.79	0.19	1.44	0.55	1.29	3.58	0.35	0.31	0.15
Control	00.0 -/+	4/- 0.07	+/- 0.16	+/- 0.21	+/- 0:15	+/- 0.03	+/- 0.07	+/- 0.14	+/- 0.71	+/- 0.11	+/- 0.39	00.0 -/+	+/- 0.01	+/- 0.02	+/- 0.01	+/- 0.04	90.0 -/+	+/- 0.29	+/- 0.05	00.0 -/+	00.0 -/+
Con	0.28	2.22	1.80	1.19	2.86	0.72	1.17	2.03	14.67	4.01	3.81	0.56	49.46	0.26	1.34	0.48	1.32	2.77	0.37	0.29	0.21
	FasL	ក្នុ	Faim	Daxx	FADD	Caspase-1	Caspase-2	Caspase-3	Caspase-6	Caspase-7	Caspase-8	Caspase-11	Cytochrome C	Apaf-1	DFF45	DFF40	Bad	Bax	Bcl-2L	Bcl-2ª	Ptpn13

15

20

10

Ŋ

References

5

10

15

- 1. Semsei I. (2000) On the nature of aging. Mech Aging Dev 117:93-108.
- 2. Sohal, RS, Weindruch, R. (1998) Oxidative stress, caloric restriction, and aging. Science 273:59-63.
- 3. Finch, CE, Revkun, G. (2001) The genetics of aging. Annu. Rev. Genom. Hum. Genet. 2:435-462.
- 4. Roth, GS, Lasnikov, V, Lesnikov, M, Ingram, DK, Land, MA (2001) Dietary caloric restriction prevents the age-related decline in plasma melatonin levels of rhesus monkeys. J Clin Endocrinol Metab. 86:3292-5.
- 5. Roth GS, Lane MA, Ingram DK, Mattison JA, Elahi D, Tobin JD, Muller D, Metter EJ (2002) Biomarkers of caloric restriction may predict longevity in humans. Science. 297:811-813.
- 6. Walford RL, Mock D, Verdery R, MacCallum T. (2002) Calorie restriction in biosphere 2: alterations in physiologic, hematologic, hormonal, and biochemical parameters in humans restricted for a 2-year period. J Gerontol A Biol Sci Med Sci 57:211-24.
- 7. Kenyon C, Chang J, Gensch E, Rudner A, Tabtiang R. (1993) A C. elegans mutant that lives twice as long as wild type. Nature 366:461-464.
- 8. Lin, K, Dorman, JB, Rodan, A, Kenyon, C. (1997). daf-16: 25 an HNF-3/Forkhead family member that can function to double the life-span of Caenorhabditis elegans. Science 278, 1319-1322.
 - 9. Clancy DJ, Gems D, Harshman LG, Oldham S, Stocker H, Hafen

- 12. Ramaswamy S, Nakamura N, Sansal I, Bergeron L, Sellers WR. (2002) A novel mechanism of gene regulation and tumor suppression by the transcription factor FKHR. Cancer Cell 2002 2:81-91.
- 5 13. Hekimi, S, Guarente, L. (2003) Genetics and the specificity of the aging process. Science 299:1351-1354.
 - 14. Brown-Borg, HM, Borg, KE, Meliska, CJ, Bartke, A. (1996) Dwarf mice and the aging process. Nature 384:33.
 - 15. Flurkey K, Papaconstantinou J, Miller RA, Harrison DE.
- 10 (2001) Lifespan extension and delayed immune and collagen aging in mutant mice with defects in growth hormone production.

 Proc Natl Acad Sci USA 98:6736-6741.
 - 16. Zhou, Y, Xu, BC, Maheshwari, HG, He, L, Reed, M, Lozykowski, M, Okada, S, Cataldo, L, Coschigano, K, Wagner, TE,
- Baumann, G, Kopchick, JJ. (1997) A mammalian model for Laron syndrome produced by targeted disruption of the mouse growth hormone receptor/binding protein gene (the Laron mouse). Proc. Nat. Acad. Sci. USA 94:13215-13220.
- 17. Coschigano, K, Clemmons, D, Bellush, LL, Kopchick, JJ.

 (2000) Assessment of growth parameters and life-span of GHR/BP
 gene-disrupted mice. Endocrinology 141:2608-2613.
- 17a. Coschigano, KT, Holland, AN, Riders, ME, List, EO, Flyvberg, A, Kopchick, JJ, Deletion, but not antagonism, of the mouse growth hormone receptor results in severely decreased body weights, insulin and IGF-1 levels and increased lifespan, Endocrinology (electronically published May 30, 2003 as doi:10.1210/en.2003-0374).
- 18. Holzenberger M, Dupont J, Ducos B, Leneuve P, Geloen A, Even PC, Cervera P, Le Bouc Y. (2003) IGF-1 receptor regulates lifespan and resistance to oxidative stress in mice.

 Nature 421:182-187.
- 19. Migliaccio E, Giorgio M, Mele S, Pelicci G, Reboldi P, Pandolfi PP, Lanfrancone L, Pelicci PG. (1999) The p66shc

445

adaptor protein controls oxidative stress response and life span in mammals. Nature 402:309-313.

- 20. Bartke A, Wright JC, Mattison JA, Ingram DK, Miller RA, Roth GS. (2001) Extending the lifespan of long-lived mice. Nature 414:412.
- 21. Weindruch R, Kayo T, Lee CK, Prolla TA. (2002) Gene expression profiling of aging using DNA microarrays. Mech Aging Dev 123:177-193.
- 22. Lee CK, Allison DB, Brand J, Weindruch R, Prolla TA.

 (2002) Transcriptional profiles associated with aging and middle age-onset caloric restriction in mouse hearts. Proc Natl Acad Sci USA 99:14988-14993.
 - 23. Prolla TA. (2002) DNA microarray analysis of the aging brain. Chem Senses 27299-306.

446

Citation of documents herein is not intended as an admission that any of the documents cited herein is pertinent prior art, or an admission that the cited documents is considered material to the patentability of any of the claims of the present application. All statements as to the date or representation as to the contents of these documents is based on the information available to the applicant and does not constitute any admission as to the correctness of the dates or contents of these documents.

The appended claims are to be treated as a non-limiting recitation of preferred embodiments.

10

15

20

25

30

35

In addition to those set forth elsewhere, the following references are hereby incorporated by reference, in their most recent editions as of the time of filing of this application: Kay, Phage Display of Peptides and Proteins: A Laboratory Manual; the John Wiley and Sons Current Protocols series, including Ausubel, Current Protocols in Molecular Biology; Coligan, Current Protocols in Protein Science; Coligan, Current Protocols in Immunology; Current Protocols in Human Genetics; Current Protocols in Cytometry; Current Protocols Pharmacology; Current Protocols in Neuroscience; Current Protocols in Cell Biology; Current Protocols in Toxicology; Current Protocols in Field Analytical Chemistry; Current Protocols in Nucleic Acid Chemistry; and Current Protocols in and the following Cold Spring Harbor Human Genetics; Laboratory publications: Sambrook, Molecular Cloning: A Laboratory Manual; Harlow, Antibodies: A Laboratory Manual; Manipulating the Mouse Embryo: A Laboratory Manual; Methods in Yeast Genetics: A Cold Spring Harbor Laboratory Course Manual; Drosophila Protocols; Imaqing Neurons: A Laboratory Early Development of Xenopus laevis: A Laboratory Manual; Manual; Using Antibodies: A Laboratory Manual; At the Bench: A Laboratory Navigator; Cells: A Laboratory Manual; in Yeast Genetics: A Laboratory Course Manual; Discovering Neurons: The Experimental Basis of Neuroscience; Genome

447

Analysis: A Laboratory Manual Series; Laboratory DNA Science; Strategies for Protein Purification and Characterization: A Laboratory Course Manual; Genetic Analysis of Pathogenic Bacteria: A Laboratory Manual; PCR Primer: A Laboratory Manual; Methods in Plant Molecular Biology: A Laboratory Course Manual; Manipulating the Mouse Embryo: A Laboratory Manual; Molecular Probes of the Nervous System; Experiments with Fission Yeast: A Laboratory Course Manual; A Short Course in Bacterial Genetics: A Laboratory Manual and Handbook for Escherichia coli and Related Bacteria; DNA Science: A First Course in Recombinant DNA Technology; Methods in Yeast Genetics: A Laboratory Course Manual; Molecular Biology of Plants: A Laboratory Course Manual.

5

10

. 15

20

25

30

35

All references cited herein, including journal articles or abstracts, published, corresponding, prior or otherwise related U.S. or foreign patent applications, issued U.S. or foreign patents, or any other references, are entirely incorporated by reference herein, including all data, tables, figures, and text presented in the cited references. Additionally, the entire contents of the references cited within the references cited herein are also entirely incorporated by reference.

Reference to known method steps, conventional methods steps, known methods or conventional methods is not in any way an admission that any aspect, description or embodiment of the present invention is disclosed, taught or suggested in the relevant art.

The foregoing description of the specific embodiments will so fully reveal the general nature of the invention that others can, by applying knowledge within the skill of the art (including the contents of the references cited herein), readily modify and/or adapt for various applications such specific embodiments, without undue experimentation, without departing from the general concept of the present invention. Therefore, such adaptations and modifications are intended to

448

be within the meaning and range of equivalents of the disclosed embodiments, based on the teaching and guidance presented herein. It is to be understood that the phraseology or terminology herein is for the purpose of description and not of limitation, such that the terminology or phraseology of the present specification is to be interpreted by the skilled artisan in light of the teachings and guidance presented herein, in combination with the knowledge of one of ordinary skill in the art.

5

10

15

20

25

30

Any description of a class or range as being useful or preferred in the practice of the invention shall be deemed a description of any subclass (e.g., a disclosed class with one or more disclosed members omitted) or subrange contained therein, as well as a separate description of each individual member or value in said class or range.

The description of preferred embodiments individually shall be deemed a description of any possible combination of such preferred embodiments, except for combinations which are impossible (e.g, mutually exclusive choices for an element of the invention) or which are expressly excluded by this specification.

If an embodiment of this invention is disclosed in the prior art, the description of the invention shall be deemed to include the invention as herein disclosed with such embodiment excised.

PCT/US2004/017322

CLAIMS

.5

1. A method of (I) reducing a rate of biological aging in a human subject, and/or(II) delaying the time of onset, or reducing the severity, of an undesirable age-related phenotype, and/or (III) protecting against an age-related (senescent) disease, which comprises

administering to the subject a protective amount of an agent which is

(1) a polypeptide which is substantially structurally identical 10 or conservatively identical in sequence to a reference protein which is (a) selected from the group consisting of mouse and human proteins set forth in master table 1, subtable 1A, or (b) selected from the group consisting of human proteins within at least one of the human protein classes set forth in master 15 table 2, subtable 2A,

or

- (2) an expression vector encoding the polypeptide of (1) above 20 and expressible in a human cell, under conditions conducive to expression of the polypeptide of (1);
- where said agent reduces a rate of biological aging in said subject, and/or delays the time of onset, or reduces the 25 severity, of an undesirable age-related phenotype in said subject, and/or protects against an age-related disease.
- 2. A method of (I) reducing a rate of biological aging in a human subject, and/or(II) delaying the time of onset, or 30 reducing the severity, of an undesirable age-related phenotype, and/or (III) protecting against an age-related (senescent) disease, which comprises administering to the subject a protective amount of an agent which is

450

- (1) an antagonist of a polypeptide, occurring in said subject, which is substantially structurally identical or conservatively identical in sequence to a reference protein which is (a) selected from the group consisting of mouse and human proteins set forth in master table 1, subtable 1B, or (b) selected from the group consisting of human proteins belonging to at least one of the human protein classes set forth in master table 2, subtable 2B,
- 10 (2) an anti-sense vector which inhibits expression of said polypeptide in said subject,

5-

15

20

25

30

35

where said agent reduces a rate of biological aging in said subject, and/or delays the time of onset, or reduces the severity, of an undesirable age-related phenotype in said subject, and/or protects against an age-related disease.

3. A method of determining a biological age of a human subject, or a rate of biological aging of a human subject, which comprises

assaying tissue or body fluid samples from said subjects to determine the level of expression of a "favorable" human marker gene, said human marker gene encoding a human protein which is substantially structurally identical or conservatively identical in sequence to a reference protein which is (a) selected from the group consisting of mouse and human proteins set forth in master table 1, subtable 1A, or (b) selected from the group consisting of human proteins within at least one of the human protein classes set forth in master table 2, subtable 2A,

and inversely correlating the level of expression of said marker gene with a biological age or a rate of biological aging of said patient.

451

- 4. A method of determining a biological age of a human subject, or a rate of biological aging of a human subject, which comprises
- assaying tissue or body fluid samples from said subjects to 5 determine the level of expression of an "unfavorable" marker gene, said human marker gene encoding a human protein identical substantially structurally which is conservatively identical in sequence to a reference protein which is (a) selected from the group consisting of mouse and 10 human proteins set forth in master table 1, subtable 1B, or (b) selected from the group consisting of human proteins belonging to at least one of the human protein classes set forth in master table 2, subtable 2B,

and directly correlating the level of expression of said marker gene with a biological age or a rate of biological aging of said subject.

5. The method of claims 1 or 2 in which (I) applies.

- 6. The method of claims 1 or 2 in which (II) applies.
- 7. The method of claims 1 or 2 in which (III) applies.
- 8. The method of claim 5 in which biological age is measured by a biomarker.
- 9. The method of claim 8 in which the marker is a simple 30 biomarker.
 - 10. The method of claim 8 in which the marker is a composite biomarker.
- 11. The method of claim 5 in which the affected biological age is the overall biological age of the subject.

452

- 12. The method of claim 5 in which the affected biological age is the biological age of a body system of the subject.
- 13. The method of claim 5 in which the affected biological age is the biological age of an organ of the subject.
 - 14. The method of claim 13 in which the organ is the liver.
- 15. The method of claim 8 in which at least one marker is the level of a biochemical in the blood of the subject.
 - 16. The method of claim 15 in which the biochemical is growth hormone or IGF-1.
- 15 17. The method of any one of claims 1-16 in which (a) applies.

20

- 18. The method of any one of claims 1-17 in which the reference protein is a human protein.
- 19. The method of any one of claims 1-17 in which the reference protein is a mouse protein.
- 20. The method of any one of claims 3 or 4 in which the level of expression of the marker protein is ascertained by measuring the level of the corresponding messenger RNA.
- 21. The method of any one of claims 3 or 4in which the level of expression is ascertained by measuring the level of a protein encoded by said marker gene.
 - 22. The method of any one of claims 1-21 in which said polypeptide is at least 80% identical or at least highly conservatively identical to said reference protein.
 - 23. The method of any one of claims 1-22 in which said

453

polypeptide is at least 90% identical to said reference protein.

- 24. The method of claim 23 in which said polypeptide is identical to said reference protein.
 - 25. The method of any one of claims 1-24 in which the E-value cited for the reference protein in Master Table 1 is not more than e-6.

26. The method of claim 25 in which the E-value cited for the reference protein in Master Table 1 is less than e-10.

- 27. The method of claim 26 in which the E value calculated by BLASTN or BLASTX is than e-15, more preferably less than e-20, still more preferably less than e-40, even more preferably less than e-60, considerably more preferably less than e-80, and most preferably less than e-100.
- 28. The method of claims 2 or 4, or of any of claims 5-27 to the extent dependent on 2 or 4, in which the antagonist is an antibody, or an antigen-specific binding fragment of an antibody.
- 25 29. The method of claims 2 or 4, or of any of claims 5-27 to the extent dependent on 2 or 4, in which the antagonist is a peptide, peptioid, nucleic acid, or peptide nucleic acid oligomer.
- 30. The method of claims 2 or 4, or of any of claims 5-27 to the extent dependent on 2 or 4, in which the antagonist is an organic molecule with a molecular weight of less than 500 daltons.
- 35 31. The method of claim 30 in which said organic molecule is identifiable as a molecule which binds said polypeptide by

454 screening a combinatorial library.

32. The method of any one of claims 2 or 4, or of any of claims 5-31 to the extent dependent on 2 or 4, in which the marker protein is CIDE-A.

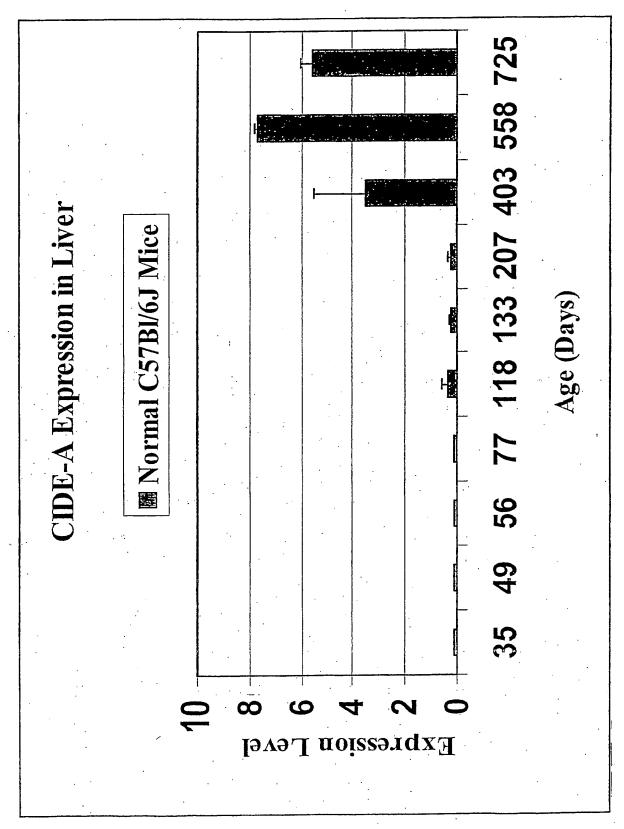


FIGURE 1

